

**ALGORITHMIC DETERMINATION OF FLANKING DNA SEQUENCES  
THAT CONTROL THE EXPRESSION OF SETS OF GENES IN  
PROKARYOTIC, ARCHEA AND EUKARYOTIC GENOMES**

**5       Reference to Related Application**

The present application is the subject of Provisional Application Serial No. 60/208,650 filed June 2, 2000 entitled ALGORITHMIC DETERMINATION OF CONNECTRONS FOR THE HIGH LEVEL REGULATION OF GENE EXPRESSION.

**10       Introduction**

RNA introduced into a cell by a virus is now known to trigger a cellular defense mechanism known as post-transcriptional gene silencing (PTGS). If the viral RNA sequence matches a sequence within the cell's genome the associated genes are turned off or silenced. This phenomenon is also called 'RNA interference' or RNAi. A single-stranded RNA can interact with another single-stranded RNA (known as antisense RNA). The single-stranded RNA can also form a triple-stranded complex with double-stranded DNA. This triple-stranded complex is known as a Hoogsteen helix. This patent application shows how two specific adjacent RNA single-stranded sequences (called C1 and C2 – for Control Sequence 1 and Control Sequence 2) interact with two distant double-stranded DNA sequences (called T1 and T2 – for Target Sequence 1 and Target Sequence 2) to form a tetradic relationship which is called a "connectron". The two distant DNA double-stranded sequences (T1 and T2) must be on the same chromosome in a genome and they must be between about 1kb and 105kb of each other. The adjacent single-stranded RNA sequences (C1/C2) can be on the same or different chromosome as the T1 and T2 sequences. The C1 sequence is identical to the T1 sequence and the C2 sequence is identical to the T2

sequence. The connectron acts to stabilize the double-stranded DNA by allowing 30nm chromatin particles to form. Genes that lie between the T1 and T2 sequences when wrapped up in 30nm chromatin particles are not open to promotion and expression. The connectron (i.e. the tetradic relationship between the T1-T2 sequences and C1/C2 sequences) provides a general explanation for PTGS. A connectron can be implemented by RNA sequences, PNA (Peptide Nucleic Acid) sequences or by a zinc-finger DNA Binding Protein (DBP) specific to the T1 and T2 sequences.

Characteristically the adjacent C1/C2 sequences lie in the 3'UTR of a gene. The T1 and T2 sequences do not lie within the translated region of any gene. These sequences "surround" one or more genes. There are, however, T1 and T2 sequence pairs that surround one or more C1/C2 sequences that are not 3'UTR to any gene. These are called "geneless connectrons". There may be promoter sequences that cause the transcription of these 3'UTR sequences.

A computer-based algorithm that is similar to the algorithm used in the US Patent 6,205,404 has been developed to determine the connectron structure of any genome. This algorithm determines the existence of all the connectrons in the genomic DNA. Connectrons exist in prokaryotes, archaea, single-celled eukaryotes, multi-celled eukaryotes, plants and higher animals. Connectron relationships exist between prokaryotes and their plasmids. The geneless connectrons provide a possible mechanism for forming a hierarchy of gene expression control that will produce an understanding of cell differentiation and tissue development.

Each connectron is a unique tetrad of sequences. Each connectron changes the expression of the genes between the T1 and T2 sequences. The C1 sequence (which is equivalent to the T1 sequence) and the C2 sequence (which is equivalent to the T2 sequence) are determined by the invention described in this patent application. In general, the tetrad of connectron sequences can be patented because the structure of matter is known and the function of specific gene expression modulation is also known. Gene expression modification can be produced by introducing antisense

RNA or PNA to interact C1/C2 RNA sequences or zinc-finger DBPs to interact with the T1 and T2 sequences. Using connectrons it will be possible to modify cellular and tissue behavior in a very general manner.

Examples will be given from different genomes to illustrate that the connectron is a perfectly general and universal concept.

### Definitions

Double stranded DNA – Watson and Crick showed in 1953 that DNA naturally forms a double-stranded helix. A typical double stranded sequence is

5'-TAGAGGAGTACCAC-3'

3'-ATCTCCTCATGGTG-5'

Hydrogen Bond - The force between a hydrogen atom and another heavier atom such as Oxygen (O), Nitrogen (N), Phosphorus (P), or Sulfur (S).

Positive strand – The positive strand is normally represented 5' to 3' running left to right as in

5'-TAGAGGAGTACCAC-3'

Negative strand – The negative strand is normally represented 5' to 3' running right to left as in

3'-ATCTCCTCATGGTG-5'

Single stranded RNA – Either the positive or the negative strand of the double-stranded DNA can be transcribed by the polymerase. In RNA U replaces T.

5 RNA of positive strand sequence 5'-UAGAGGAGUACCAC-3'  
RNA of negative strand sequence 5'-GUGGUACUCCUCUA-3'

Antisense RNA – The antisense strand of any RNA sequence is the complement sequence

10

RNA sequence 5'-UAGAGGAGUACCAC-3'  
Antisense RNA sequence 3'-AUCUCCUCAUGGUG-5'

Triple Strand Helix – The RNA sequence of a RNA/DNA triple-strand complex is the same as the positive strand of the DNA

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DNA positive strand 5'-TAGAGGAGTACCAC-3'  
DNA negative strand 3'-ATCTCCTCATGGTG-5'  
RNA strand 5'-UAGAGGAGUACCAC-3'

20

Promoter – Any region of DNA, that binds proteins which engage the polymerase transcription mechanism.

TATA Box – A region near the 3' end of a promoter with the sequence TATA.

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mRNA – The RNA produced from the DNA by the polymerase as a result of transcription

Start of transcription – The 3' end of a promoter where the polymerase mechanism begins to transcribe DNA into mRNA.

30

Exon – Any region of mRNA which is used to code for proteins



Intron – Any region of mRNA lying between two exons which is not used to code for proteins. The introns are edited out of the initial RNA transcript to form the mature mRNA.

5

3' UTR – The untranslated 3' end of an mRNA is beyond the end of the last exon. A stop codon in the mRNA causes the ribosome to stop the translation of mRNA into protein.

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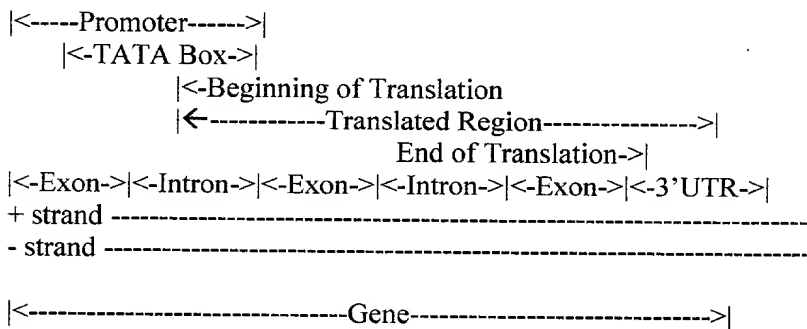
End of translation – The 3' end of the 3'-most exon.

Translated region – Any collection of exons and introns.

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Gene – Any DNA region that codes for a protein. Introns do not occur in prokaryotic genes and they sometime fail to occur in eukaryotic genes. A typical model of a gene is

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Positive strand gene – Any gene in which the features run 5' to 3' on the positive strand

Negative strand gene – Any gene in which the features run 5' to 3' on the negative strand

35

C1 sequence – Any positive or negative strand DNA sequence of 20 bases or more.

The C2 sequence must occur in the same chromosome as the C1 sequence.

C2 sequence – Any positive or negative strand DNA sequence of 20 bases or more.

The C1 sequence must occur in the same chromosome as the C2 sequence.

5

C1/C2 – Any positive or negative strand DNA sequence of 40 or more bases such that the C1 sequence is adjacent to the C2 sequence

10

T1 sequence – Any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence. The T1 and T2 sequences must be between about 1kb and 105kb apart.

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T2 sequence – Any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence. The T2 and T1 sequences must be between about 1kb and 105kb apart.

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Last exon gap or Gap-Distance – The number of bases between the end of transcription and the beginning of the C1/C2 sequence. In prokaryotes and single-celled eukaryotes this gap can range from no bases to 500 bases. In multi-celled eukaryotes the gap can be as large as 10,000 bases.

Poly-adenylation signal – A number of Adenosine (A) bases are added to the mRNA at the end of the 3'UTR.

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Possible Connectron – Any set of T1, T2 and C1/C2 sequences such that the C1 sequence is identical to the T1 sequence and the C2 sequence is identical to the T2 sequence. The promoter of some gene causes the mRNA of the gene to be expressed. The mRNA is edited to eliminate the introns. The whole mRNA including the 3'UTR can move about in the cell or the nucleus of the cell. The C1/C2 RNA that is part of the 3'UTR moves to the T1 and T2 DNA sequences. A triple-stranded complex of the DNA and the RNA forms such that the C1 sequence forms hydrogen bonds with the T1 sequence and the C2 sequence forms hydrogen bonds with the T2 sequence.

30

Because the C1 sequence is adjacent to the C2 sequence, the T1 sequence is brought physically close to the T2 sequence. This produces a loop of between about 1kb and 105kb in the DNA. Histone proteins reduce the length of the DNA by binding 200 bases. Histone/DNA complexes form six-fold symmetry chromatin assemblies. The diameter of the chromatin assemblies is approximately 30nm.

Real Connectron – Any Possible Connectron which is within the Gap-Distance of some gene

Homologous connectron – The T1 sequence and the T2 sequence are on the same chromosome as the C1/C2 sequence

Heterologous connectron – The T1 sequence and the T2 sequence are on a chromosome different from chromosome of the C1/C2 sequence

Permanent connectron – Any C1/C2 sequence, which is 3' UTR to some gene that is not surrounded by any T1 and T2 sequence pairs

Transient connectron – Any C1/C2 sequence, which is 3' UTR to some gene that is surrounded by one or more T1 and T2 sequence pairs

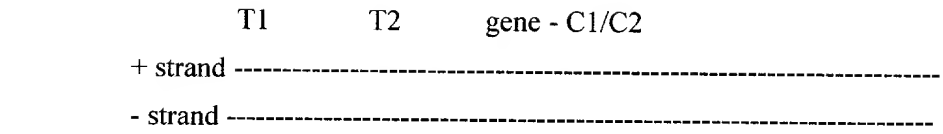
Self-limiting connectron – Any C1/C2 sequence which is 3'UTR to some gene that is surrounded by the T1 and T2 sequences such that C1=T1 and C2=T2

Geneless connectron - Any C1/C2 sequence which is not 3'UTR to some gene but is surrounded by some T1 and T2. A promoter may lie 5' to the C1/C2 sequence.

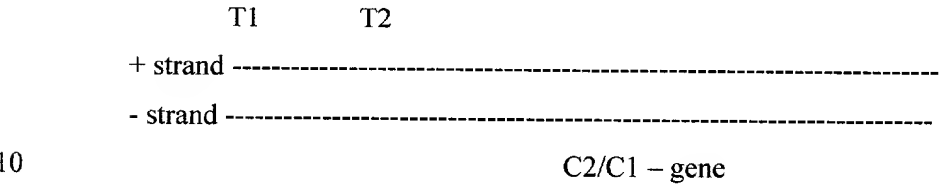
Bidirectionality of Connectron Excitation – A C1/C2 short loop on one strand selects a T1-T2 long loop pair on the same or the opposite strand. The C1/C2 short loop has a complementary C1'/C2' sequence on the opposite strand. Similarly the T1-T2 long loop pair has a complementary long loop pair T1'-T2'. Wherever a C1/C2, T1-T2 tetrad exists there is a complementary C1'/C2', T1'-T2' tetrad. The C1/C2 short loop

can be transcribed as a 3'UTR to a gene on the same strand. The C1'/C2' short loop which is on the strand opposite to the C1/C2 short loop can also can be transcribed as a 3'UTR to a gene on the same strand. There are four possible models of action

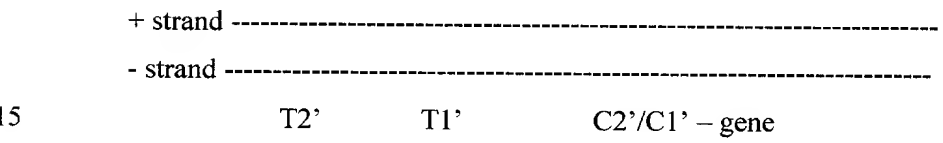
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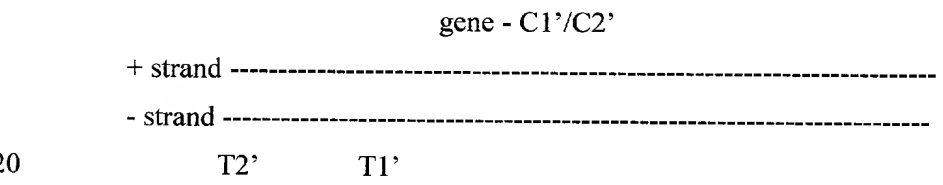
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Of course, the short loops and the long loops do not have to be on the same chromosome.

25 Hierarchy of connectron action – When a C1/C2 is expressed it forms a T1-T2 loop by forming a connectron. The C1/C2 sequence does not have to be on the same chromosome as the T1 and T2 sequences. This provides a way of causing interaction between chromosomes. When the T1-T2 loop forms, any genes in that loop region which had been expressing C1/C2 sequences in their 3'UTRs, now cease expressing the C1/C2 sequences. The connectrons formed by these C1/C2 sequences will cease to exist after some time thus opening up the genes inside the respective T1-T2 loops

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to expression. The hierarchy of connectron action is alternates between repression and expression. The connectron hierarchies can be of any depth.

5 One-to-Many connectron action – One C1/C2 sequence can form connectrons in many different places on many different chromosomes. The only requirement is that C1=T1 and C2=T2. This makes it possible for one expression event to control the expression of many genes on different chromosomes.

10 Many-to-One connectron action – C1/C2s that come from many different places on many different chromosomes can form a connectron for a specific T1-T2 sequence pair. The only requirement is that C1=T1 and C2=T2. This makes it possible for many different expression events to control the expression of one set of genes on a particular chromosome.

15 Many-to-Many connectron action – The arrangement of C1/C2s and T1-T2s across chromosomes can form a complex web of gene expression control relationships.

20 Percentage of the Genome Regulated by Connectrons – Since the connectrons for a sequenced genome can be calculated, the percentage of the genome that is open to connectron regulation can be known.

25 Emergent Property – The network of connectrons in any genome emerges from a knowledge of the complete DNA sequence of the genome. Because both the C1/C2 sequences and the T1-T2 sequences can be any place in the genome, the whole genomic sequence must be known before all the connectrons can be determined.

30 Paradigm Shift – For the past fifty years since the discovery by Watson and Crick of the double-helical nature of DNA, the reigning paradigm for scientific discovery has been the study of one gene and its effects on the behavior of a cell. The advent of genomic sequencing and this invention of connectrons that emerge from the whole genome will produce a shift in the way scientists view biological systems and the way they formulate and execute experiments. The many-to-many relationships between

the connectrons means that there are many ways in which the expression of a set of genes can be modulated. The multiplicity of control pathways means produces a system stability that makes it possible for biological systems to be stable for long periods of evolutionary time. The thinking that goes into formulating scientific experiments will have to change to accommodate the changes in understanding that will be induced by the application and extension of this patent application.

Hierarchy of DNA Structuring – The DNA of a cell's genome is structured in a hierarchy of six levels. Figures 1, 2 and 3 have been adapted from The Molecular Biology of the Cell by Alberts, Bray, Lewis, Raff, Roberts and Watson [third edition pages 354, 345 and 348]. As shown in figure 1, the double stranded DNA is level 1. The double-stranded DNA is wrapped around histone proteins to form a chromatin particle that is level 2 of the hierarchy. Level 2 is described as “beads-on-a-string” in figure 1. The chromatin particles are packed in a six-fold symmetry as shown in figure 2a and figure 2b. These six-fold assemblies have a diameter of 30 nm. Each 30 nm assembly contains from 18 (i.e.  $6 * 3$ ) to 30 (i.e.  $6 * 5$ ) chromatin particles. The 30 nm assemblies aggregate into large loops which range in length from 5,000 bases to 100,000 bases of DNA. The size of these large loops as shown in figure 1 is approximately 300 nm. These large loops constitute level 4 of the structuring hierarchy. As shown in figure 1, level 5 of the DNA structuring hierarchy many large loops are condensed to form a structure which is approximately 700 nm in diameter. The complete chromosome that constitutes level 6 of the hierarchy is composed of two very long sections of level 5 DNA.

Model of Chromatin Structure – The level 4 structure of DNA as shown in figure 1 ranges in length from 5,000 to 105,000 bases of DNA. Figure 3 shows that proteins are thought to connect portions of the long loops formed by the 30 nm particles to form a chromosome axis. These condensed long loops are described as chromomeres in The Molecular Biology of the Cell.

## Prior Art

The chromomere model of DNA structuring was presented by N. A Resnik, et al.[1] and is based on electron microscopic data. There are more recent papers studying a variety of genomes with electron microscopy but no equivalent study of chromomeres has been done on a fully sequenced genome.

A recent News Feature in Nature by T. Gura [2] described the discovery of post-transcriptional gene silencing in which viral RNA interacts with the transcribed RNA of the cell to silence the expression of genes. This article describes experiments in *C. elegans* and *D. megalomaster* in which RNA that is complementary to mRNA introduced into a cell. This "antisense" RNA has the effect of turning off the expression of one or more genes. The introduced complementary RNA produces an "RNA interference" called RNAi.

Thomas Werner and his colleagues at Genomatix in Munich, Germany have developed an approach to understanding what they call "Matrix Attachment Region" (MAR). Figure 5 shows their interpretation of the structure of DNA surrounding a gene. The following description of the MAR is copied from the Genomatix web site

"Matrix Attachment Regions (MARs) MARs are sequence regions that are responsible for the attachment of genomic DNA to the nuclear matrix or scaffold. Transcription absolutely requires anchorage of genomic DNA to the nuclear matrix.

### Functional features of MARs:

Anchoring of regulatory elements like promoters and enhancers to the nuclear matrix.

Ensuring long term activity of promoters and enhancers in chromatin.





## Brief Description of the Objects of the Invention

5 An object of the invention is to provide a method of identifying DNA sequences that control the expression of different collections of genes in a genome comprising, detecting selected DNA sequences adjacent to some genes excluding exons and introns.

10 An object of the invention is to provide a method of identifying DNA sequences that control the expression of different collections of genes comprising, detecting, by computer, one or more pairs of non-adjacent DNA sequences to which are bound to two RNA sequences.

15 An object of the invention is to provide a method of identifying DNA sequences that control the expression of different collections of genes in a genome comprising detecting changes in connectron behavior in the genome.

20 An object of the invention is to provide a method of modifying the expression of different gene collections in a genome, comprising detecting changes in connectron behavior as a result of an exogenous stimulus.

25 An object of the invention is to provide a method of detecting where and when new genes are being integrated into a host genome comprising detecting the connectrons in said host genome.

An object of the invention is to provide a method of detecting the expression effect of different gene collections in a given body comprising detecting the back and forth flow of connectrons between the chromosomes thereof.

An object of the invention is to provide a method of modifying a given body comprising modifying the connectron organization therein.

An object of the invention is to provide a method of detecting connectron control and target sequences in a given genome comprising:

determining the base composition of said genome,  
determining one or more sites of control sequence organization, and/or  
determining one or more sites of target application.

An object of the invention is to provide a method of determining the response of a cell in any tissue to changes in the cell's environment and/or genetic composition comprising providing a complete genomic DNA sequence for the organism and determining the effect of changes in connectrons due to application of a given exogenous stimulus to the genome.

An object of the invention is to provide a method of determining in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes, the tetradic relationship  $T1=C1$  and  $T2=C2$  where T1 and T2 are DNA sequences 20 or more bases in length, where the C1 sequence is adjacent to the C2 sequence, where the T1 and T2 sequences are on the same chromosome, and where the C1/C2 sequences are on the same chromosome as T1 and T2 or where the C1/C2 sequences are on a chromosome different from T1 and T2, wherein:

C1 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C2 sequence must occur in the same chromosome as the C1 sequence,

C2 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C1 sequence must occur in the same chromosome as the C2 sequence,

C1/C2 - any positive or negative strand DNA sequence of 40 or more bases such that the C1 sequence is adjacent to the C2 sequence,

T1 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence, the T1 and T2 sequences must be between about 1kb and 105kb apart, and

T2 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence, the T2 or T1 sequences must be between about 1kb and 105kb apart.

An object of the invention is to provide a method of determining in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes, the connectron relationship that permits many different C1/C2 short loops to control the existence of a T1-T2 long loop and wherein said C1/C2 short lops can be on the same chromosome or on different chromosomes from the T1-T2 long loop, wherein:

C1 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C2 sequence must occur in the same chromosome as the C1 sequence,

C2 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C1 sequence must occur in the same chromosome as the C2 sequence,

C1/C2 - any positive or negative strand DNA sequence of 40 or more bases such that the C1 sequence is adjacent to the C2 sequence,

T1 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence, the T1 and T2 sequences must be between about 1kb and 105kb apart, and

T2 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence, the T2 or T1 sequences must be between about 1kb and 105kb apart.

5 An object of the invention is to provide a method of determining in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes, the connectron relationship that permits one C1/C2 short loop to control the existence of many T1-T2 long loops, the C1/C2 short loop can be on the same chromosome or on different chromosomes from the T1-T2 long loops, wherein:

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C1 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C2 sequence must occur in the same chromosome as the C1 sequence,

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C2 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C1 sequence must occur in the same chromosome as the C2 sequence,

20

C1/C2 - any positive or negative strand DNA sequence of 40 or more bases such that the C1 sequence is adjacent to the C2 sequence,

25

T1 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence, the T1 and T2 sequences must be between about 1kb and 105kb apart, and

T2 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence, the T2 or T1 sequences must be between about 1kb and 105kb apart.

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An object of the invention is to provide a method of determining in the connectron relationships between prokaryotes and their plasmids wherein said connectrons implement a control mechanism between the two genomes that makes it possible from

them to form a symbiotic relationship, and in the case of *D. radiodurans* the relationship is not symmetric, and the *D. radiodurans* genome sends C1/C2 short loops to the MP1 plasmid, wherein:

5 C1 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C2 sequence must occur in the same chromosome as the C1 sequence,

10 C2 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C1 sequence must occur in the same chromosome as the C2 sequence,

15 C1/C2 - any positive or negative strand DNA sequence of 40 or more bases such that the C1 sequence is adjacent to the C2 sequence,

20 T1 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence, the T1 and T2 sequences must be between about 1kb and 105kb apart, and

25 T2 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence, the T2 or T1 sequences must be between about 1kb and 105kb apart.

30 An object of the invention is to provide a method of determining that connectron relationships that exist in plant and higher animals.

An object of the invention is to provide a method of determining in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes, the connectron relationship that permits one C1/C2 short loop to control the existence of one or more T1-T2 long loops without being subject to any expression controls other than those of the gene to which the C1/C2 is 3'UTR, wherein:



C2 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C1 sequence must occur in the same chromosome as the C2 sequence,

5

C1/C2 - any positive or negative strand DNA sequence of 540 or more bases such that the C1 sequence is adjacent to the C2 sequence,

10

T1 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence, the T1 and T2 sequences must be between about 1kb and 105kb apart, and

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T2 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence, the T2 or T1 sequences must be between about 1kb and 105kb apart.

An object of the invention is to provide a method of determining in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes, the connectron relationship that permits one C1/C2 short loop to control the existence of the T1-T2 long loop that surrounds it, wherein:

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C1 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C2 sequence must occur in the same chromosome as the C1 sequence,

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C2 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C1 sequence must occur in the same chromosome as the C2 sequence,

30

C1/C2 - any positive or negative strand DNA sequence of 40 or more bases such that the C1 sequence is adjacent to the C2 sequence,



T1 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence, the T1 and T2 sequences must be between about 1kb and 105kb apart, and

5 T2 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence, the T2 or T1 sequences must be between about 1kb and 105kb apart.

10 An object of the invention is to provide a method of determining the connectron relationships that do not have any genes within the T1-T2 long loop, wherein:

T1 sequence is any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T2 sequence, and

15 T2 sequence - any positive or negative strand DNA sequence of 20 bases or more that is on the same chromosome as the T1 sequence, and the T2 or T1 sequences must be between about 1kb and 105kb apart.

20 An object of the invention is to provide a method of determining the geneless connectron relationship where one C1/C2 short loop controls the existence of many geneless T1-T2 long loops, wherein:

25 C1 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C2 sequence must occur in the same chromosome as the C1 sequence,

30 C2 sequence - any positive or negative strand DNA sequence of 20 bases or more, the C1 sequence must occur in the same chromosome as the C2 sequence,

C1/C2 - any positive or negative strand DNA sequence of 40 or more bases such that the C1 sequence is adjacent to the C2 sequence,



## Description of the Drawings and Tables

The above and other objects, advantages and features of the invention will become more apparent when considered with the following specification and accompanying drawings and tables wherein:

Figure 1 DNA is structured in six levels of increasing condensation. Double stranded DNA is level 1. Two turns of DNA are wrapped about each chromatin particle at level 2. The chromatin particles which each containing 200 base pairs form into 30 nm particles at level 3. The 30 nm particles form into large loops with an approximate dimension of 300 nm at level 4. Metaphase chromosomes form a condensed structure with an approximate dimension of 700 nm at level 5. An entire metaphase chromosome has a width of approximately 1400 nm at level 6. The large loops at level 4 of the DNA structuring are thought to have between 20,000 (20 kb) and 100,000 (100 kb) base pairs.

The Molecular Biology of the Cell by Alberts, Bray, Lewis, Raff, Roberts and Watson, 3rd. ed. , Garland Publishing, Inc., New York, 1994, p. 354

Figure 2 (a) Chromatin DNA forms into a six-fold symmetry 30nm particles.  
(b) The six-fold symmetry 30nm particles form a linear chain with a varying number of repeat units.

The Molecular Biology of the Cell by Alberts, Bray, Lewis, Raff, Roberts and Watson , 3rd. ed. , Garland Publishing, Inc., New York, 1994, p. 345

Figure 3	Long loops of 30nm particles are thought to be closed at the bottom of the loop by proteins.
5	The Molecular Biology of the Cell by Alberts, Bray, Lewis, Raff, Roberts and Watson, 3rd. ed. , Garland Publishing, Inc., New York, 1994, p. 348
Figure 4	(a) Transcription and Editing. (b) Movement of the RNA through the Nucleus. (c) Connectron Formation
Figure 5	Overview of schematic organization of a typical transcriptionally active chromosomal loop.
10	From <a href="http://genomatix.gsf.de/func_genomics/functional_genomics.html">http://genomatix.gsf.de/func_genomics/functional_genomics.html</a>
Table 1	Connectron Properties for Prokaryotic, Archea and Eukaryotic Genomes
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Table 2	Yeast Inter-Chromosomal Connectron Distribution

Figure 6 Genome size plotted as a log-log function of the Number of Connectrons

Figure 7 Number of Sequence Instances plotted as a function of the Number of Fragments

5 Figure 8 Level 0 – The overall view of the algorithm

Figure 9 Level 1 – Process Flow of the Algorithm

Figure 10 Level 2a – two pages – Process Genome into Blocking Fragment File

Figure 11 Level 2b – two pages – Compute the Connectrons for a Genome

Figure 12 Level 2c – two pages – Analyze Possible Connectrons

10 Figure 13 Level 3a – Setup Genome Usage Memory

Figure 14 Level 3b – Find DBP-Size Blocking File for T1-Window

Figure 15 Level 1 – Find DBP-Size Blocking File for T2-Window

Figure 16 Level 2a – two pages – Find C1/C2 Entries

Figure 17 Level 2b – two pages – Scan Genome Usage Memory for Potential Connectrons

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## Description of the Invention

A connectron is a relationship among four DNA sequences. Each sequence must be at least 20 bases long. There is a report by Sharp and Zamore [3] that RNA sequences of “about length 25” are important as sources of RNAi. 27 bases were actually used as the minimum length of each of the sequences. The T1 sequence is on one strand of some chromosome in a genome. The T2 sequence is on the same strand of the same chromosome as the T1 sequence. The T1 and T2 sequences (which are each at least 20 bases in length) must be at least 5,000 bases distant from each other but they can not be more than 105,000 bases distant from each other. The C1 sequence and the C2 sequence (which are each at least 20 bases in length) are adjacent to each other on some strand of some chromosome in the genome. The C1/C2 sequences – called the “short loop” – can be on the same strand as the T1 and T2 sequences or they can be on the opposite strand. The C1/C2 sequences of the short loop can be on the same chromosome as the T1 and T2 sequences but they can also be on a different chromosome in the genome. When a genome has only one chromosome, then the point is moot. Many genomes, of course, have several chromosomes. The C1 sequence is identical to the T1 sequence and the C2 sequence is identical to the T2 sequence.

The C1/C2 sequence must be on the same strand as a gene, either be directly adjacent to the gene (i.e. a gap of 0 bases) for prokaryotic genomes or at this time be within 10,000 bases for eukaryotic genomes. The size of the gap between the end of the gene and the beginning of the C1/C2 sequence is a variable. The C1/C2 short loop is expressed as the 3’UTR (Un-Translated Region) of the gene. In the case of prokaryotic genes that do not normally have introns, the whole mRNA becomes the active species for connectron formation. In the case of eukaryotic genes, the whole transcript is the active species for connectron formation upon editing of the transcript to eliminate the introns. The whole transcript then can move about in the cytoplasm of prokaryotic cells or the nucleus of eukaryotic cells. Since the C1 sequence is equivalent to the T1 sequence and the C2 sequence is equivalent to the T2 sequence,

the C1 RNA can form a Hoogsteen triple-stranded RNA/DNA/DNA helix with the double-stranded T1 sequence. Similarly the C2 RNA can form a Hoogsteen triple-stranded RNA/DNA/DNA helix with the double-stranded T2 sequence. Because the C1 sequence and the C2 sequence are adjacent to each other, the C1/T2 RNA/DNA/DNA Hoogsteen triple helix is brought into physical adjacency to the C2/T2 RNA/DNA/DNA Hoogsteen triple helix. RNA/DNA/DNA hybrid helices are the most stable form of triple helix. RNA double helices, DNA double helices, RNA triple helices and DNA triple helices are all significantly less stable than a RNA/double-stranded DNA triple helix. The stable physical adjacency of the two triple-stranded Hoogsteen helices ensures that the long loop of double-stranded DNA between the T1 sequence and the T2 sequence can then be structured into 30 nm chromatin particles as shown in level 4 of figure 1. The genes on either strand of the DNA between the T1 sequence and the T2 sequence when they are structured into the 30 nm chromatin particles are not open to promotion and expression.

The tetradic relationship between the T1 and T2 sequences that form the long loop and the C1/C2 sequences that form the short loop are called a connectron. The name “connectron” was suggested by J. David Rawn Ph.D. of Towson University. A connectron is possible if the T1, T2, C1 and C2 sequences exist. A connectron is real if the C1/C2 short loop sequence is adjacent to an expressible gene. If the expression of the adjacent gene is inside one or more T1 – T2 long loops then this connectron is said to be transient. If the adjacent gene is not inside any possible T1-T2 long loop then the connectron is said to be permanent. If a connectron is inside of a T1-T2 long loop that has the same sequences (i.e. T1 is really equal to C1 and T2 is really equal to C2) then the connectron is said to be self-limiting. This is true because once the C1/C2 sequence is expressed it forms the T1-T2 long loop that then shuts off the expression of the gene adjacent to the C1/C2 sequence. Self-limiting connectrons can also be called “spike” connectrons since they generate a short-duration spike of the C1/C2 short loop sequence. If a T1-T2 long loop does not contain any genes but it contains C1/C2 short loop sequences then this type of connectrons is said to be geneless. The C1/C2 short loops within a geneless T1-T2 long loop can, of course, control the expression of genes.

The physical existence and lifetimes of the connectrons must be proved by molecular biological experimentation. This physical experimental process, however, is logically quite separate from the computational experimentation that have been conducted from June of 1999 to May of 2001. The computational search for the existence of connectrons has been extremely positive. These computations have shown that connectrons exist in prokaryotes, in archaea, between prokaryotes and their plasmids, in single-celled eukaryotes, in multi-celled eukaryotes, in plants, in higher animals and in humans. All of these features and properties are described in the claims section that follows.

The connectron invention is very powerful. It depends only on sequence equivalency. The minimum length of the four sequences seems to be about 20 bases. In the calculations shown in this patent application, 27 bases have been used as a minimum. The Nature News Feature [1] says that other scientists have found RNA sequences of length about 25 that have interesting gene silencing properties. The Nature article does not give any mechanism. Because of my algorithm and its use on a variety of genomes, this patent application provides the computational proof that a particular mechanism is highly probable. The connectron invention provides an explanation for how communication occurs with a chromosome as well as between chromosomes in genomes that have more than one chromosome. Since each T1-T2 long loop can contain one or more genes, the connectron invention provides a mechanism for turning on and turning off sets of genes simultaneously. In time, the connectron invention will provide an explanation for how differentiation of how one cell's behavior differs from the behavior of another adjacent cell. It is already clear from the computational experiments that have been made on *S. cerevisiae*, *C. elegans* and *D. megalomaster* that the number of geneless connectrons increases dramatically as evolution proceeds from single-celled eukaryotes (i.e. *S. cerevisiae*) to 1,000 cell eukaryotes (i.e. *C. elegans*) to visible creatures (i.e. *D. megalomaster*). The extension of this evolutionary progress to plants (i.e. *A. thaliana*) for which only three chromosomes are sequenced and humans (i.e. *H. sapiens*) for which only one chromosome is completely sequenced. Although the complete human genome was



published in Nature and Science in February of 2001, the NIH-sponsored genomic sequencing results are available for about 1/3 of the bases in the whole genome. The human genomic sequence determined by Celera Genomics, Inc. is available only by subscription. Table 1 shows how the genome size, the number of genes, the number of gene-containing and geneless connectrons and the percentage of genes controlled are related in many different genomes.

The C1/C2 short loops originate on one chromosome. The T1-T2 long loops can be on the same or different chromosomes. Table 2 which is for yeast (*S. cerevisiae*) is a square matrix of how many C1/C2 short loops on a given chromosome are sent to form T1-T2 long loops on other chromosomes. The diagonal of this matrix shows that many chromosomes send connectrons to themselves. The striking feature of this particular table is that chromosome 6 only sends connectrons to chromosome 12 but that it receives connectrons from chromosomes 4,5,7,10,12,13,15 and 16.

Any tetrad of connectron sequences (i.e. the T1, T2, C1 and C2 sequences) as well as the fact of the adjacency of the C1/C2 short loop sequence to the transcribing gene can be patented because the content of matter and the utility can be exactly described. The utility of a connectron is that the T1-T2 long loop shuts off the expression of the genes that lie between the T1 sequence and the T2 sequence. In the case of geneless connectrons, the utility is of a higher level in that the C1/C2 short loops contained in the higher-level geneless T1-T2 long loop, eventually form other lower-level T1-T2 long loops around a set of genes.

The invention of connectrons comes at a particularly important time in biological discovery. The geneless connectrons make a many-to-many hierarchical control mechanism possible. It is already clear from the determination of the connectrons for *C. elegans* and *D. megalomaster* that there are as many or more geneless connectrons than there are genes. It has been clear for some time that the number of genes in a genome is not particularly correlated with the size of the genome. Figure 6 shows that the size of a genome is roughly linearly correlated with the number of connectrons.



## Detailed Description of the Invention

The algorithm for the determination of connectrons in any genome or any genome fragment is represented in the following flow diagrams. The Level 0 diagram in figure 8 shows the general relationships in a digital computer. The central processor of the digital computer uses the computer program to take genome descriptors, the genomic DNA sequences and the tables of gene features to produce a file of blocking fragments and a file of the optimal connectrons for the genome. The printer serves to make hard copies of the files and this patent application. The level 1 diagram in figure 9 shows the three essential steps in the determination of connectrons. The genome is first processed into a blocking fragment file. Then the blocking fragments are used to compute the connectrons for the genome. Finally the potential connectrons are analyzed to determine if the C1/C2 sequences are in the 3'UTR of a gene. The level 2a diagram in figure 10 shows the steps required for the processing of the genome into a file of blocking fragments. The genomic DNA sequence is decomposed into 27-base frames for both the positive and negative strands. These fragments are written to the unsorted fragment file. The fragment file is then sorted is then read and formed into groups of equivalent sequences. The (.blk) file contains the sequence and a pointer to the (.gptr) file which contains the pointers to the position of the fragments in the genomes. The position in the genome includes the chromosome number, the position in the chromosome and the strand (i.e. positive and negative). A sample of these files follows

Sample of the (.blk) file for *S. cerevisiae*

27-base fragment	Number of instances	Pointer to (.gptr) file
------------------	------------------------	----------------------------

11111111111111111111111111111111	0	1
111111123244233313332443414	1	2
111111141113443133314333341	2	4
111111232442333133324434141	1	5
111111323311133323144423444	2	7
111111332213331341414443413	2	9
111111333444112343412323243	1	10

	111111333444113343412323243	9	19
	111111411134431333143333414	2	21
	111111443223134142124434124	2	23
	111112223234344444443144442	2	25
5	111112244123441122214421213	8	33
	111112311241114344334134431	2	35
	111112324423331333244341414	1	36
	111112344232231344242234342	1	37
	111112433444244421144134211	1	38
10	111112444311313442332142224	1	39
	111113131241131114424413231	1	40
	111113143332344311113133411	1	41
	111113233111333231444234441	2	43

15 In fragments above 1=G, 2=C, 3=A, 4=T

Sample of the (.gptr) file for *S. cerevisiae*

There are 16 chromosomes in *S. cerevisiae*

20	Item	Chromosome	Position in Chromosome	Direction
	1	0	0	0
25	2	4	11137	1
	3	12	467619	1
	4	12	458482	1
	5	4	11138	1
	6	12	465759	2
30	7	12	456622	1
	8	1	219366	1
	9	8	539978	1
	10	14	522451	1
	11	4	1099073	1
35	12	4	1210003	1
	13	7	539068	1
	14	12	654136	1
	15	12	596455	1
	16	15	121016	1
40	17	15	598127	2
	18	16	847724	1
	19	16	59765	1
	20	12	467620	1
	21	12	458483	1
45	22	12	461657	1
	23	12	452520	1
	24	13	838006	1

	25	15	288270	1
	26	4	83593	1
	27	4	992867	1
5	28	6	162265	1
	29	7	845687	1
	30	10	531560	2
	31	15	282208	1
	32	16	860418	1
10	33	16	572308	1
	34	12	465992	1
	35	12	456855	1
	36	4	11139	1
	37	8	89343	1
15	38	4	10302	1
	39	1	19894	2
	40	16	9311	1
	41	10	735203	1
	42	12	465760	1
20	43	12	456623	1

In direction column above 1=positive strand, 2=negative strand

The level 2b diagram in figure 11 shows the computation of the connectrons. The genome descriptors consist of the number and length of the chromosomes. The algorithm uses an array that represents several facts about each base position in the genome. The level 3a diagram in figure 13 shows the setup of the Genome-Usage memory. The gene features are used to prevent the region of the genome that codes for proteins from being used for the connectron sequences (i.e. the T1s, the T2s, the C1s and the C2s). In the level 2a diagram of figure 10, the algorithm steps through each chromosome and within each chromosome through each base position looking for acceptable T1-windows of 27 bases. A T1-window can be used to form a connectron relationship if there are two or more instances of this fragment in the blocking fragment file. The computation in the level 3b diagram of figure 14 determines if the T1-window is acceptable or not. Once an acceptable T1-window is found, the algorithm (in the level 2a diagram of figure 10) looks for acceptable T2-window positions that lie between 5,000 and 105,000 bases from the T1-window. The computation for determining acceptable T2-window positions is done in the level 3c diagram of figure 15. Once a pair of T1 and T2 window positions are found, the



## Examples

The algorithm for the determination of optimal connectrons has been applied to a number of different publicly available genomes. The connectron is a tetradic relationship between four sequence elements – T1, T2, C1 and C2. The claims presented in this section are written by the program NearGene that implements the flow diagram Level 2c of figure 12. The examples are written a uniform type of English. Each example contains some or all of the following elements

-----

10        Name of genome  
          Description of T1  
          Length of T1-T2 loop  
          The chromosome on which the T1-T2 loop exists  
          The identifier number within the genome of the T1 sequence  
15        The T1 sequence  
          Description of T2  
          The identifier number within the genome of the T2 sequence  
          The T2 sequence  
          A list of genes whose expression is controlled by the T1-T2 loop  
20        The common names of the genes as obtained from the NCBI gene feature file  
          (.ptt)  
          A list of C1/C2 short loops whose expression if controlled by the T1-T2 loop  
          The chromosome on which the C1/C2 short loop exists  
          The common name of the gene which expresses the C1/C2 short loop as an  
25        RNA  
          The sequence of the C1/C2 short loop  
          A list of C1/C2 short loops that control the formation of the T1-T2 loop  
          The chromosome on which the C1/C2 short loop exists  
          The common name of the gene which expresses the C1/C2 short loop as an  
30        RNA  
          The sequence of the C1/C2 short loop

The match between the C1/C2 sequence and the T2 sequence

[illegible]

5

10

Table 1. Demographic characteristics of the study population	
Age (years)	65.0 ± 10.0
Gender	
Male	50 (50.0%)
Female	50 (50.0%)
Education (years)	12.0 ± 2.0
Marital status	
Married	40 (80.0%)
Single	10 (20.0%)
Occupation	
Retired	30 (60.0%)
Unemployed	20 (40.0%)
Income (USD/month)	1,200 ± 300
Health status	
Good	30 (60.0%)
Fair	20 (40.0%)
Poor	10 (20.0%)
Comorbidities	
Hypertension	20 (40.0%)
Diabetes	10 (20.0%)
Cholesterol	15 (30.0%)
Arthritis	10 (20.0%)
Depression	5 (10.0%)
Other	5 (10.0%)
Medication	
Yes	30 (60.0%)
No	20 (40.0%)
Medication type	
Antidepressants	10 (20.0%)
Antipsychotics	5 (10.0%)
Mood stabilizers	5 (10.0%)
Other	10 (20.0%)
Duration of illness (years)	5.0 ± 3.0
Family history	
Yes	10 (20.0%)
No	40 (80.0%)
Family size	3.0 ± 1.0
Living with family	40 (80.0%)
Living alone	10 (20.0%)
Religious beliefs	
Religious	30 (60.0%)
Non-religious	20 (40.0%)
Religious practices	
Regular	20 (40.0%)
Irregular	10 (20.0%)
None	20 (40.0%)
Religious beliefs impact	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on health	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on medication	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on lifestyle	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on social support	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on coping strategies	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on self-management	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on adherence	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on quality of life	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)
Religious beliefs impact on overall health	
High	10 (20.0%)
Medium	20 (40.0%)
Low	20 (40.0%)



## Index of Pages for Connectron Samples

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- 5      1.      Connectrons occur in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes.

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- 10      2.      Many Connectrons control the expression of one set of genes in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes.

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- 15      3.      One connectron controls the expression of many sets of genes in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes.

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- 20      4.      Connectrons occur between prokaryotes and their plasmids.

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- 25      5.      Connectrons occur in plants and higher animals

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- 30      6.      Permanent connectrons exist in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes.

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- 35      7.      Transient connectrons exist in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes.

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- 40      8.      Self-limiting connectrons occur in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes

5

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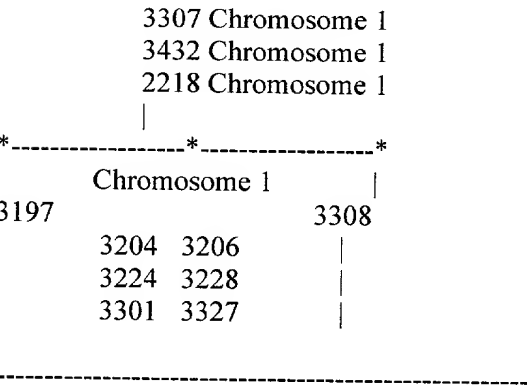
- | Table 1. Demographic characteristics of the study population |                |
|--|----------------|
| Age (years)  | Mean (SD)      |
| 18-24  | 20.5 (2.5)     |
| 25-34  | 29.5 (4.5)     |
| 35-44  | 39.5 (5.5)     |
| 45-54  | 49.5 (6.5)     |
| 55-64  | 59.5 (7.5)     |
| 65-74  | 69.5 (8.5)     |
| 75-84  | 79.5 (9.5)     |
| 85-94  | 89.5 (10.5)    |
| 95-104   | 99.5 (11.5)    |
| 105-114  | 109.5 (12.5)   |
| 115-124  | 119.5 (13.5)   |
| 125-134  | 129.5 (14.5)   |
| 135-144  | 139.5 (15.5)   |
| 145-154  | 149.5 (16.5)   |
| 155-164  | 159.5 (17.5)   |
| 165-174  | 169.5 (18.5)   |
| 175-184  | 179.5 (19.5)   |
| 185-194  | 189.5 (20.5)   |
| 195-204  | 199.5 (21.5)   |
| 205-214  | 209.5 (22.5)   |
| 215-224  | 219.5 (23.5)   |
| 225-234  | 229.5 (24.5)   |
| 235-244  | 239.5 (25.5)   |
| 245-254  | 249.5 (26.5)   |
| 255-264  | 259.5 (27.5)   |
| 265-274  | 269.5 (28.5)   |
| 275-284  | 279.5 (29.5)   |
| 285-294  | 289.5 (30.5)   |
| 295-304  | 299.5 (31.5)   |
| 305-314  | 309.5 (32.5)   |
| 315-324  | 319.5 (33.5)   |
| 325-334  | 329.5 (34.5)   |
| 335-344  | 339.5 (35.5)   |
| 345-354  | 349.5 (36.5)   |
| 355-364  | 359.5 (37.5)   |
| 365-374  | 369.5 (38.5)   |
| 375-384  | 379.5 (39.5)   |
| 385-394  | 389.5 (40.5)   |
| 395-404  | 399.5 (41.5)   |
| 405-414  | 409.5 (42.5)   |
| 415-424  | 419.5 (43.5)   |
| 425-434  | 429.5 (44.5)   |
| 435-444  | 439.5 (45.5)   |
| 445-454  | 449.5 (46.5)   |
| 455-464  | 459.5 (47.5)   |
| 465-474  | 469.5 (48.5)   |
| 475-484  | 479.5 (49.5)   |
| 485-494  | 489.5 (50.5)   |
| 495-504  | 499.5 (51.5)   |
| 505-514  | 509.5 (52.5)   |
| 515-524  | 519.5 (53.5)   |
| 525-534  | 529.5 (54.5)   |
| 535-544  | 539.5 (55.5)   |
| 545-554  | 549.5 (56.5)   |
| 555-564  | 559.5 (57.5)   |
| 565-574  | 569.5 (58.5)   |
| 575-584  | 579.5 (59.5)   |
| 585-594  | 589.5 (60.5)   |
| 595-604  | 599.5 (61.5)   |
| 605-614  | 609.5 (62.5)   |
| 615-624  | 619.5 (63.5)   |
| 625-634  | 629.5 (64.5)   |
| 635-644  | 639.5 (65.5)   |
| 645-654  | 649.5 (66.5)   |
| 655-664  | 659.5 (67.5)   |
| 665-674  | 669.5 (68.5)   |
| 675-684  | 679.5 (69.5)   |
| 685-694  | 689.5 (70.5)   |
| 695-704  | 699.5 (71.5)   |
| 705-714  | 709.5 (72.5)   |
| 715-724  | 719.5 (73.5)   |
| 725-734  | 729.5 (74.5)   |
| 735-744  | 739.5 (75.5)   |
| 745-754  | 749.5 (76.5)   |
| 755-764  | 759.5 (77.5)   |
| 765-774  | 769.5 (78.5)   |
| 775-784  | 779.5 (79.5)   |
| 785-794  | 789.5 (80.5)   |
| 795-804  | 799.5 (81.5)   |
| 805-814  | 809.5 (82.5)   |
| 815-824  | 819.5 (83.5)   |
| 825-834  | 829.5 (84.5)   |
| 835-844  | 839.5 (85.5)   |
| 845-854  | 849.5 (86.5)   |
| 855-864  | 859.5 (87.5)   |
| 865-874  | 869.5 (88.5)   |
| 875-884  | 879.5 (89.5)   |
| 885-894  | 889.5 (90.5)   |
| 895-904  | 899.5 (91.5)   |
| 905-914  | 909.5 (92.5)   |
| 915-924  | 919.5 (93.5)   |
| 925-934  | 929.5 (94.5)   |
| 935-944  | 939.5 (95.5)   |
| 945-954  | 949.5 (96.5)   |
| 955-964  | 959.5 (97.5)   |
| 965-974  | 969.5 (98.5)   |
| 975-984  | 979.5 (99.5)   |
| 985-994  | 989.5 (100.5)  |
| 995-1004   | 999.5 (101.5)  |
| 1005-1014  | 1009.5 (102.5) |
| 1015-1024  | 1019.5 (103.5) |
| 1025-1034  | 1029.5 (104.5) |
| 1035-1044  | 1039.5 (105.5) |
| 1045-1054  | 1049.5 (106.5) |
| 1055-1064  | 1059.5 (107.5) |
| 1065-1074  | 1069.5 (108.5) |
| 1075-1084  | 1079.5 (109.5) |
| 1085-1094  | 1089.5 (110.5) |
| 1095-1104  | 1099.5 (111.5) |
| 1105-1114  | 1109.5 (112.5) |
| 1115-1124  | 1119.5 (113.5) |
| 1125-1134  |                |

1. Connectrons occur in prokaryotes, archae, single-celled eukaryotes and multi-celled eukaryotes.

Connectrons exist as tetradic relationships where the sequence T1 is equivalent to the sequence C1 (written T1=C1) and where the sequence T2 equals the sequence C2 (written T2=C2) where T1 and T2 are DNA sequences 20 or more bases in length, where the C1 sequence is adjacent to the C2 sequence, where the T1 and T2 sequences are on the same chromosome, and where the C1/C2 sequences are on the same chromosome as T1 and T2 or where the C1/C2 sequences are on a chromosome different from T1 and T2. The connectron relationship has been found to exist in prokaryotes, archae, single-celled eukaryotes and multi-celled eukaryotes.

Example of a prokaryote connectron – E. coli

In this example the existence of the T1-T2 (3197-3308) long loop is controlled by three C1/C2 short loops (3307, 3432 and 2218). The T1-T2 long loop controls the expression of 64 genes on chromosome 1 in addition to six C1/C2 (3204, 3206, 3223, 3228, 3301 and 3327) short loops. The C1/C2 short loop 3327 lies outside the range of the T1-T2 long loop (3197-3308) but this C1/C2 is expressed as a 3'UTR to the gene hemG that is within the range of the T1-T2 long loop.



Connectron control elements for chromosome 1 of the E. coli genome

A double stranded DNA loop of length 93.542 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3197. This T1 control element has the DNA sequence

AAAAAATGCGCGGTCAGAAAATTATTTTAAATTTCTCTTGTTCAGGCCGG  
AATAACTCCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACG  
CCGCCGGGTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAA  
ATAAATGCTTGACTCTGTAGCGGGAA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 3308. This T2 control element has the DNA sequence

TAAATTTCTCTTGTTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTG  
ACACGGAACAACGGCAAACACGCCGCCGGGTCAGCGGGGTTCTCCTGAG  
AACTCCGGCAGAGAAAGCAAAAATAAATGCTTGACTCTGTAGCGGGAAG  
GCGTATTATGCACACCCCGCGCCGCT

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

rrsC	gltU	rrlC	rrfC	aspT	trpT	yifA	yifE	yifB
ilvL	ilvG_1	ilvM	ilvE	ilvD	ilvA	ilvY	ilvC	ppiC
b3776	rep	gppA	rhlB	trxA	rhoL	rho	rfe	wzzE
wecB	rffH	wecD	wecE	wzxE	yifM_2	wecG	yifK	
argX	hisR	leuT	proM	aslB	aslA	hemY	hemX	
hemD	cyaA	cyaY	b3808	dapF	uvrD	b3814	corA	
yigF	yigG	rarD	yigI	pldA	recQ	yigJ	yigK	pldB
yigL	yigM	metR	metE	ysgA	udp	yigN	ubiE	yigP

b3836    yigU    yigW\_1    rfaH    yigC    ubiB    fadA    fadB  
pepQ    trkH    hemG

This long T1/T2 double stranded DNA loop modulates the expression of the  
following C1/C2 short loops

A C1/C2 short loop on chromosome 1 whose identifier is 3204 controls the  
expression of the genes of one or more other T1/T2 long loops. This C1/C2 short  
loop is expressed as a RNA single strand that is 3'UTR to the gene rrsC and has the  
DNA sequence

GATGTGCCCAGATGGGATTAGCTAGTAGGTGGGGTAACGGCTCACCTAGG  
CGACGATCCCTAGCTGGTCTGAGAGGATGACCAGCCACACTGGAACTGAG  
ACACGGTCCAGACTCCTACGGGAGGCAGCAGTGGGGAATATTGCACAATG  
GGCGCAAGCCTGATGCAGCCATGCCGCGTGTATGAA

A C1/C2 short loop on chromosome 1 whose identifier is 3206 controls the  
expression of the genes of one or more other T1/T2 long loops. This C1/C2 short  
loop is expressed as a RNA single strand that is 3'UTR to the gene rrsC and has the  
DNA sequence

GTCCCCTTCGTCTAGAGGCCAGGACACCGCCCTTTCACGGCGGTAACAG  
GGGTTCGAATCCCCTAGGGGACGCCACTTGCTGGTTTGTGAGTGAAAGTC  
ACCTGCCTTAATATCTCAAACTCATCTTCGGGTGATGTTTGAGATATTTG  
CTCTTTAAAAATCTGGATCAAGCTGAAAATTGAAA

A C1/C2 short loop on chromosome 1 whose identifier is 3223 controls the  
expression of the genes of one or more other T1/T2 long loops. This C1/C2 short  
loop is expressed as a RNA single strand that is 3'UTR to the gene rrlC and has the  
DNA sequence

GCTGAAGTAGGTCCCAAGGGTATGGCTGTTCGCCATTTAAAGTGGTACGC  
 GAGCTGGGTTTAGAACGTCGTGAGACAGTTCGGTCCCTATCTGCCGTGGG  
 CGCTGGAGAACTGAGGGGGGCTGCTCCTAGTACGAGAGGACCGGAGTGG  
 ACGCATCACTGGTGTTCGGGTTGTCATGCCAATGGCA

5

A C1/C2 short loop on chromosome 1 whose identifier is 3225 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene *rrlC* and has the DNA sequence

10

AAACAGAATTTGCCTGGCGGCCGTAGCGCGGTGGTCCCACCTGACCCCAT  
 GCCGAACTCAGAAGTGAAACGCCGTAGCGCCGATGGTAGTGTGGGGTCTC  
 CCCATGCGAGAGTAGGGAAGTCCAGGCATCAAATTAAGCAGTA

15

A C1/C2 short loop on chromosome 1 whose identifier is 3228 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene *rrfC* and has the DNA sequence

20

GGTCATAAAACCGGTGGTTGTAAAAGAATTCGGTGGAGCGGTAGTTCAGT  
 CGGTTAGAATACCTGCCTGTCACGCAGGGGGTCGCGGGTTCGAGTCCCGT  
 CCGTTCCGCCAC

25

A C1/C2 short loop on chromosome 1 whose identifier is 3301 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene *ubiB* and has the DNA sequence

30

TTATCGTGCCTACAAATAGTCCGAACCGTAGGCCGGATAAGGCGTTTACG  
 CCGCATC

A C1/C2 short loop on chromosome 1 whose identifier is 3307 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene fadA and has the DNA sequence

5

TGCCGGATGCGGCGTAAACGCCTTATCCGGCCTACGGTTCGGACTATTTGT  
AGGCA

10 A C1/C2 short loop on chromosome 1 whose identifier is 3327 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene hemG and has the DNA sequence

15 AAAAAATGCGCGGTCAGAAAATTATTTTAAATTTCTCTTGTCAGGCCGG  
AATAACTCCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACG  
CCGCCGGGTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAA  
ATAAATGCTTGACTCTGTAGCGGGAAGGCGTATTATG...CCCGTCACACCA  
TGGGAGTGGGTTGCAAAGAAGTAGGTAGCTTAACCTTCGGGAGGGCGCT  
20 TACCACTTTGTGATTCATGACTGGGGTGAAGTCGTAACAAGGTAACCGTA  
GGGGAACCTGCGGTTGGATCACCTCCTTACCTTAAAGAAGCGTTCTTTG

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

25 A C1/C2 short loop on chromosome 1 whose identifier is 3307 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene hemG and has the DNA sequence

30 AAAAAATGCGCGGTCAGAAAATTATTTTAAATTTCTCTTGTCAGGCCGG  
AATAACTCCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACG  
CCGCCGGGTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAA  
ATAAATGCTTGACTCTGTAGCGGGAAGGCGTATTATG...CCCGTCACACCA

TGGGAGTGGGTTGCAAAAGAAGTAGGTAGCTTAACCTTCGGGAGGGCGCT  
TACCACTTTGTGATTCATGACTGGGGTGAAGTCGTAACAAGGTAACCGTA  
GGGGAACCTGCGGTTGGATCACCTCCTTACCTTAAAGAAGCGTTCTTTG

5 The match between the T1 sequence and the C1/C2 sequence is

AAAAAATGCGCGGTCAGAAAATTATTTTAAATTCCTCTTGTCAGGCCGG  
AATAACTCCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACG  
CCGCCGGGTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAA  
10 ATAAATGCTTGACTCTGTAGCGGAA

The match between the T2 sequence and the C1/C2 sequence is

TAAATTCCTCTTGTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTG  
15 ACACGGAACAACGGCAAACACGCCCGCGGGTCAGCGGGGTTCTCCTGAG  
AACTCCGGCAGAGAAAGCAAAAATAAATGCTTGACTCTGTAGCGGGAAG  
GCGTATTATGCACACCCCGCGCCGCT

20 A C1/C2 short loop on chromosome 1 whose identifier is 3432 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene btuB and has the DNA sequence

TGCGCGGTCAGAAAATTATTTTAAATTCCTCTTGTCAGGCCGGAATAACT  
CCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACGCCGCCGG  
25 GTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAAATAAATG  
CTTGACTCTGTAGCGGGAAGGCGTATTATGCACACC...ACACCATGGGAGT  
GGGTTGCAAAAGAAGTAGGTAGCTTAACCTTCGGGAGGGCGCTTACCACT  
TTGTGATTCATGACTGGGGTGAAGTCGTAACAAGGTAACCGTAGGGGAAC  
CTGCGGTTGGATCACCTCCTTACCTTAAAGAAGCGT

30 The match between the T1 sequence and the C1/C2 sequence is



TGCGCGGTCAGAAAATTATTTTAAATTTCTCTTGTCAGGCCGGAATAACT  
 CCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACGCCGCCGG  
 GTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAAATAAATG  
 CTTGACTCTGTAGCGGGAA

5

The match between the T2 sequence and the C1/C2 sequence is

TAAATTTCTCTTGTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTG  
 ACACGGAACAACGGCAAACACGCCGCCGGGTCAGCGGGGTTCTCCTGAG  
 AACTCCGGCAGAGAAAGCAAAAATAAATGCTTGACTCTGTAGCGGGAAG  
 GCGTATTATGCACACCCCGCGCCGCT

10

A C1/C2 short loop on chromosome 1 whose identifier is 2218 controls the  
 expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
 as a RNA single strand that is 3'UTR to the gene clpB and has the DNA sequence

15

CTTGTGAGGCCGGAATAACTCCCTATAATGCGCCACCACTGACACGGAAC  
 AACGGCAAACACGCCGCCGGGC

20

The match between the T1 sequence and the C1/C2 sequence is

CTTGTGAGGCCGGAATAACTCCCTATAATGCGCCACCACTGACACGGAAC  
 AACGGCAAACACGCCGCCGGGC

25

The match between the T2 sequence and the C1/C2 sequence is

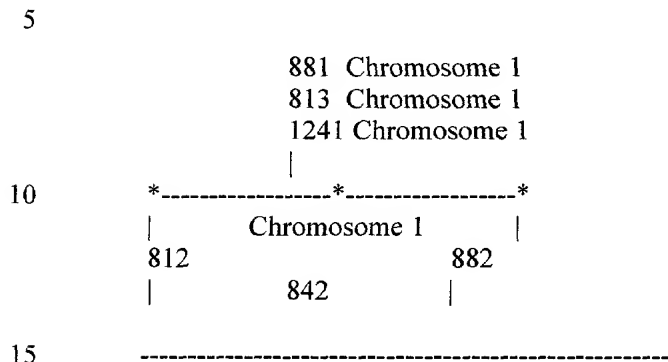
CTTGTGAGGCCGGAATAACTCCCTATAATGCGCCACCACTGACACGGAAC  
 AACGGCAAACACGCCGCCGGGTC

30

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Example of an archea connectron – *H. pylori*

In this example the existence of the T1-T2 (812-882) long loop is controlled by three C1/C2 short loops (881, 813 and 1214). The T1-T2 long loop controls the expression of 54 genes on chromosome 1 in addition to one C1/C2 (843) short loop.



Connectron control elements for chromosome 1 of *H. pylori* genome

A double stranded DNA loop of length 96.385 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 812. This T1 control element has the DNA sequence

TTTACTCATAGGGTTTTATAGTTCCTAGCGGAACTAAAGCA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 882. This T2 control element has the DNA sequence

TAGCGGAACTAAAGCATTCATCCCAAACACTAAAGATATTTGG

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

HP0999	HP1000	HP1001	HP1002	HP1003	HP1005	HP1006
HP1008	HP1009	HPtRNA-Pro	HP1010	HP1011	HP1013	HP1015
HP1017	HP1018	HP1020	HP1021	HP1022	HP1023	HP1024

HP1025    HP1027    HP1028    HP1030    HP1031    HP1033    HP1034  
 HP1038    HP1039    HP1040    HP1041    HP1042    HP1043    HP1044  
 HP1045    HP1046    HP1051    HP1052    HP1055    HP1056    HP1058  
 HP1060    HP1065    HPtRNA-Ser    HP1066    HP1067    HP1069    HP1070  
 5    HP1074    HP1075    HP1076    HP1077    HP1078    HP1079    HP1080  
 HP1081    HP1083    HP1084    HP1085    HP1088    HP1091    HP1092  
 HP1093    HP1094    HP1095    HP1096

10    This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

15    A C1/C2 short loop on chromosome 1 whose identifier is 813 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene HP0998 and has the DNA sequence

TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCATTTCATCCC  
 AAACACTAAAGATATTTGG

20    The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

25    A C1/C2 short loop on chromosome 1 whose identifier is 881 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene HP1096 and has the DNA sequence

TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCATTTCATCCC  
 AAACACTAAAGATATTTGG

30    The match between the T1 sequence and the C1/C2 sequence is

TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCA

The match between the T2 sequence and the C1/C2 sequence is

5 TAGCGGAACTAAAGCATTTCATCCCAAACACTAAAGATATTTGG

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

10 A C1/C2 short loop on chromosome 1 whose identifier is 813 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene HP0998 and has the DNA sequence

15 TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCATTTCATCCC  
AAACACTAAAGATATTTGG

A C1/C2 short loop on chromosome 1 whose identifier is 881 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene HP1096 and has the DNA sequence

20 TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCATTTCATCCC  
AAACACTAAAGATATTTGG

The match between the T1 sequence and the C1/C2 sequence is

25 TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCA

The match between the T2 sequence and the C1/C2 sequence is

30 TAGCGGAACTAAAGCATTTCATCCCAAACACTAAAGATATTTGG

A C1/C2 short loop on chromosome 1 whose identifier is 1241 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene HP1535 and has the DNA sequence

5 TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAAGTAAAGCATTTCATCCC  
AAACA

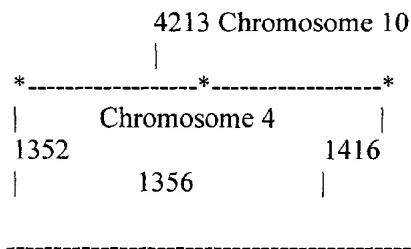
The match between the T1 sequence and the C1/C2 sequence is

TTTACTCATAGGGTTTTATAGTTCCTAGCGGAATAAGCA

The match between the T2 sequence and the C1/C2 sequence is

### Example of single-celled connectron – *S. cerevisiae*

In this example the existence of the T1-T2 (1352-1416) long loop on chromosome 4 is controlled by one C1/C2 short loop (4213) on chromosome 10. The T1-T2 long loop controls the expression of 34 genes on chromosome 4 in addition to one C1/C2 (1356) short loop.



### Connectron control elements for chromosome 1 of *S. cerevisiae* genome

A double stranded DNA loop of length 68.908 kilo-bases on chromosome 4 is bounded on the left by a T1 sequence whose identifier is 1352. This T1 control element has the DNA sequence

5

TTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1416. This T2 control element has the DNA sequence

10

ATTAGATCTATTACATTATGGGTGGTATGTTGGAATAAAAATCAACTATCA  
TCTACTAACTAGTATTTACGTTACTAGTATATTATCATATACGGTGTTAGA  
AGATGACGCAAATGATGAGAAATAGTCATCTAAATTAGTGGAAGCTGAA  
ACGCAAGGATTGATAATGTAATAGGATCAATGAATATTAACATATAAAAC  
15 GATGATAATAATATTTATAGAATTGTGTAGAATTGCAGATTCCCTTTTATG  
GATTCCTAAATCCTTGAGGAGAACTTCTAGTATATCTACATACCTAATATT  
ATAGCCTTAATCACAATGGAATCCCAACAATTACATCAAAATCCACATTC  
TCTACAGTA

20

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

25

YDR170W-A	YDR171W	YDR172W	YDR173C	YDR174W	YDR175C
YDR176W	YDR177W	YDR178W	YDR179C	YDR179W-A	YDR180W
YDR181C	YDR182W	YDR183W	YDR184C	YDR185C	YDR186C
YDR187C	YDR188W	YDR189W	YDR190C	YDR191W	YDR192C
YDR193W	YDR194C	YDR195W	YDR196C	YDR197W	YDR198C
YDR199W	YDR200C	YDR201W	YDR202C	YDR203W	YDR204W
YDR205W	YDR206W	YDR207C	YDR208W	YDR209C	YDR210W

30

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 4 whose identifier is 1356 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YDR170W-A and has the DNA sequence

AATCACACTAATCATTCTGATGATGAACTCCCTGGACACCTCCTTCTCGAT  
TCAGGAGCATCACGAACCCTTATAAGATCTGCTCATCACATACTCAGC  
ATCATCTAATCCTGACATAAACGTAGTTGATGCTCAAAAAAGAAATATAC  
CAATTAACGCTATTGGTGACCTACAATTTCACTTCCAGGACAACACCAAA  
ACATCAATAAAGGTATTGCACACTCCTAACATAGCCTATGACTTACTCAGT  
TTGAATGAATTGGCTGCAGTAGATATCACAGCATGCTTTACCAAAAACGT  
CTTAGAACG

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 10 whose identifier is 4213 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YJR029W and has the DNA sequence

ATCTATTACATTATGGGTGGTATGTTGGAATAAAAATCCACTATCGTCTAT  
CAACTAATAGTTATATTATCAATATATTATCATATACGGTGTTAAGATGAT  
GACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAAACGC  
AAGGATTGATAATGTAATAGGATCAATGAATATAAACATATAAAACGGA  
ATGAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTGA  
GGATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTGTATACCTAATA  
TTATAGCCTTTATCAACAATGGAATCCCAACAATTATCTCAACAT

The match between the T1 sequence and the C1/C2 sequence is

TTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAA

The match between the T2 sequence and the C1/C2 sequence is

5 ATCTATTACATTATGGGTGGTATGTTGGAATAAAAATC

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10

Example of a multi-celled connectron – *C. elegans*

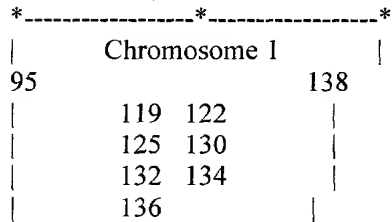
In this example the existence of the T1-T2 (9-138) long loop on chromosome 1 is controlled by three C1/C2 short loops on chromosome 5 (21719, 21949 and 21655).

15

The T1-T2 long loop controls the expression of four genes on chromosome 1 in addition to seven C1/C2 (119, 122, 125, 130, 132, 134 and 136) short loops.

20

21719 Chromosome 5  
21949 Chromosome 5  
21655 Chromosome 5



25

30

A double stranded DNA loop of length 41.978 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 95. This T1 control element has the DNA sequence

35



CAGCACGTTCTTAACCATGCAAAATCAGTTGAGAACTCTGCGTCTCTTCTC  
CCGC

5 This double stranded DNA loop is bounded on the right by a T2 control element  
whose identifier is 138. This T2 control element has the DNA sequence

ACTCTGCGTCTCTTCTCCCGCATTTTTGTAGATCA

10 This long T1/T2 double stranded DNA loop modulates the expression of the  
following genes

Y73A3A.1 Y73A3A.1 ZC123.3 ZC123.2

15 This long T1/T2 double stranded DNA loop modulates the expression of the  
following C1/C2 short loops

20 A C1/C2 short loop on chromosome 1 whose identifier is 119 controls the expression  
of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is  
expressed as a RNA single strand that is 3'UTR to the gene ZC123.3 and has the  
DNA sequence

TTGAGAACTCTGCGTCTCAACTCCCGCATTTTTGTAGATCTACGTAGATC  
AAACCGAAATGGGACACT

25 A C1/C2 short loop on chromosome 1 whose identifier is 122 controls the expression  
of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is  
expressed as a RNA single strand that is 3'UTR to the gene ZC123.3 and has the  
DNA sequence

30 GCACGGGGTTCTGGCCTTCCTCATTGAATTTTCGCGCTCCATTGACAATC  
GCCTGCCGGACAACGCGTGGGAAAGTCGTGTACTCCAC

A C1/C2 short loop on chromosome 1 whose identifier is 125 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZC123.3 and has the DNA sequence

5

ACGCGCCGTAAATCTACCCCAGATATGGCCGAGCCAAAATGGCCTAGTTC  
GGCAAACCTCTTTCATTTCAATTTATGAGGGAAGCCAGAA

10 A C1/C2 short loop on chromosome 1 whose identifier is 130 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZC123.2 and has the DNA sequence

15 CTCCCGCATTTTTTGTAGATCTACGTAGATCAAACCGAAATGAGGCACTTT  
CTGAATCCACGAGCTAGGCTTAAGCTTAGGCTTAAGCTTAGGCCTTTTCTC  
AGGCTTAGGCTTAGGCTTA

20 A C1/C2 short loop on chromosome 1 whose identifier is 132 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZC123.2 and has the DNA sequence

25 GCTTATGCTTGGGCTTAGGCTTAGGCGTAGGCTTAGGCTTAGGCTTAGGCT  
TATGCTTAGACTTAGTCTCACTATCAGTCTTAGGCTTAGGCTTAGACTTAG  
GCTTAAGCTTAGGCTTAAGCTTAGACTTAGGCTTAGGCTTAGGCTTAGGCT  
TAGGCTTAGGTTTGGGCTTAGGCTTAGGCTTAACCTC

30 A C1/C2 short loop on chromosome 1 whose identifier is 134 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZC123.2 and has the DNA sequence

TCTGCGTCTTTTCTCCCGCATTTTTTGTAGATCTACGTAGATCAAACCGAA  
 ATGAGGCACTTTCTGAATCCACGAGCTAGGCTTAAGCTTAGGCTTAAGCTT  
 AGGCCTTTTCTCAGGCTTAGGCTTAGGCTTA

5 A C1/C2 short loop on chromosome 1 whose identifier is 136 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZC123.2 and has the DNA sequence

10 GCTTATGCTTGGGCTTAGGCTTAGGCGTAGGCTTAGGCTTAGGCTTAGGCT  
 TATGCTTAGACTTAGTCTCACTATCAGTCTTAGGCTTAGGCTTAGACTTAG  
 GCTTAAGCTTAGGCTTAAGCTTAGACTTAGGCTTAGGCTTAGGCTTAGGCT  
 TAGGCTTAGGTTTGGGCTTAGGCTTAGGCTTAACCTC

15 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 5 whose identifier is 21719 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene C39F7.5 and has the DNA sequence

20 ACGTTCTTAACCATGCAAAATCAGTTGAGAACTCTGCGTCTCTTCTCCCGC  
 ATTTTTTGTAGATC

25 The match between the T1 sequence and the C1/C2 sequence is

ACGTTCTTAACCATGCAAAATCAGTTGAGAACTCTGCGTCTCTTCTCCCGC

The match between the T2 sequence and the C1/C2 sequence is

30 ACTCTGCGTCTCTTCTCCCGCATTTTTTGTAGATC

A C1/C2 short loop on chromosome 5 whose identifier is 21949 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene F16B4.4 and has the DNA sequence

5      ACCATGCAAAATCAGTTGAGAACTCTGCGTCTCTTCTCCCGCATT TTTTGT  
AGATCTACGTAGATCAAGCCGAAATGAGACACTCTGACACCACG

The match between the T1 sequence and the C1/C2 sequence is

10      ACCATGCAAAATCAGTTGAGAACTCTGCGTCTCTTCTCCCGC

The match between the T2 sequence and the C1/C2 sequence is

15      ACTCTGCGTCTCTTCTCCCGCATT TTTTGTAGATC

A C1/C2 short loop on chromosome 5 whose identifier is 21655 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene C39F7.3 and has the DNA sequence

20      AACCATGCAAAATCAGTTGAGAACTCTGCGTCTCTTCTCCCGCATT TTTTGT  
TAGATCTACG

The match between the T1 sequence and the C1/C2 sequence is

25      AACCATGCAAAATCAGTTGAGAACTCTGCGTCTCTTCTCCCGC

The match between the T2 sequence and the C1/C2 sequence is

30      ACTCTGCGTCTCTTCTCCCGCATT TTTTGTAGATC

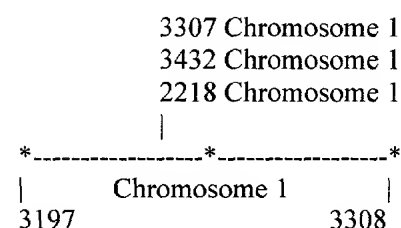
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2. Many Connectrons control the expression of one set of genes in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes.

Many different C1/C2 short loops can control the existence of one T1-T2 long loop.  
The C1/C2 short loops can be on the same chromosome or on different chromosomes from the T1-T2 long loop. This relationship is described as “many-to-one”. This relationship exists in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes

Example of a many-to-one connectron in prokaryotes – E. coli

In this example the existence of the T1-T2 (3197-3308) long loop is controlled by three C1/C2 short loops (3307, 3432 and 2218).



A double stranded DNA loop of length 93.542 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3197. This T1 control element has the DNA sequence

AAAAAATGCGCGGTCAGAAAATTATTTAAATTTCTCTTGTCAGGCCGG  
AATAACTCCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACG  
CCGCCGGGTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAA  
ATAAATGCTTGACTCTGTAGCGGGAA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 3308. This T2 control element has the DNA sequence

TAAATTTCTCTTGTTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTG  
ACACGGAACAACGGCAAACACGCCGCCGGGTCAGCGGGGTTCTCCTGAG  
AACTCCGGCAGAGAAAGCAAAAATAAATGCTTGACTCTGTAGCGGGAAG  
5 GCGTATTATGCACACCCCGCGCCGCT

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

10 rrsC    gltU    rrlC    rrfC    aspT    trpT    yifA    yifE    yifB  
ilvL    ilvG\_1    ilvM    ilvE    ilvD    ilvA    ilvY    ilvC    ppiC  
b3776    rep    gppA    rhlB    trxA    rhoL    rho    rfe    wzzE  
wecB    rffH    wecD    wecE    wzxE    yifM\_2    wecG    yifK  
argX    hisR    leuT    proM    aslB    aslA    hemY    hemX  
15 hemD    cyaA    cyaY    b3808    dapF    uvrD    b3814    corA  
yigF    yigG    rarD    yigI    pldA    recQ    yigJ    yigK    pldB  
yigL    yigM    metR    metE    ysgA    udp    yigN    ubiE    yigP  
b3836    yigU    yigW\_1    rfaH    yigC    ubiB    fadA    fadB  
pepQ    trkH    hemG

20 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

25 A C1/C2 short loop on chromosome 1 whose identifier is 3307 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene hemG and has the DNA sequence

30 AAAAAATGCGCGGTCAGAAAATTATTTTAAATTTCTCTTGTTCAGGCCGG  
AATAACTCCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACG  
CCGCCGGGTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAA  
ATAAATGCTTGACTCTGTAGCGGGAAGGCGTATTATG...GGAGTCTGCAAC  
TCGACTCCATGAAGTCGGAATCGCTAGTAATCGTGGATCAGAATGCCACG

GTGAATACGTTCCCGGGCCTTGTACACACCGCCCGTCACACCATGGGAGT  
GGGTTGCAAAAGAAGTAGGTAGCTTAACCTTCGGGAGGGGCGCTTACCACT  
TTGTGATTCATGACTGGGGTGAAGTCGTAACAAGGTAACCGTAGGGGAAC  
CTGCGGTTGGATCACCTCCTTACCTTAAAGAAGCGTTCTTTG

5

The match between the T1 sequence and the C1/C2 sequence is

AAAAAATGCGCGGTCAGAAAATTATTTTAAATTCCTCTTGTGTCAGGCCGG  
AATAACTCCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACG  
CCGCCGGGTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAA  
ATAAATGCTTGACTCTGTAGCGGGAA

10

The match between the T2 sequence and the C1/C2 sequence is

TAAATTCCTCTTGTGTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTG  
ACACGGAACAACGGCAAACACGCCCGGGTCAGCGGGGTTCTCCTGAG  
AACTCCGGCAGAGAAAGCAAAAATAAATGCTTGACTCTGTAGCGGGAAG  
GCGTATTATGCACACCCCGCGCCGCT

15

A C1/C2 short loop on chromosome 1 whose identifier is 3432 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene btuB and has the DNA sequence

20

TGCGCGGTCAGAAAATTATTTTAAATTCCTCTTGTGTCAGGCCGGAATAACT  
CCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACGCCGCCGG  
GTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAAATAAATG  
CTTGACTCTGTAGCGGGAAGGCGTATTATGCACACC...ACACCATGGGAGT  
GGGTTGCAAAAGAAGTAGGTAGCTTAACCTTCGGGAGGGGCGCTTACCACT  
TTGTGATTCATGACTGGGGTGAAGTCGTAACAAGGTAACCGTAGGGGAAC  
CTGCGGTTGGATCACCTCCTTACCTTAAAGAAGCGT

25

30

The match between the T1 sequence and the C1/C2 sequence is

5 TGC GCGGTCAGAAAATTATTTTAAATTTCTTGT CAGGCCGGAATAACT  
CCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACGCCGCCGG  
GTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAAATAAATG  
CTTGACTCTGTAGCGGGAA

The match between the T2 sequence and the C1/C2 sequence is

10 TAAATTTCTTGT CAGGCCGGAATAACTCCCTATAATGCGCCACCACTG  
ACACGGAACAACGGCAAACACGCCGCCGGGTCAGCGGGGTTCTCCTGAG  
AACTCCGGCAGAGAAAGCAAAAATAAATGCTTGACTCTGTAGCGGGAAG  
GCGTATTATGCACACCCCGCGCCGCT

15 A C1/C2 short loop on chromosome 1 whose identifier is 2218 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene clpB and has the DNA sequence

20 CTTGTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTGACACGGAAC  
AACGGCAAACACGCCGCCGGGC

The match between the T1 sequence and the C1/C2 sequence is

25 CTTGTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTGACACGGAAC  
AACGGCAAACACGCCGCCGGGC

The match between the T2 sequence and the C1/C2 sequence is

30 CTTGTCAGGCCGGAATAACTCCCTATAATGCGCCACCACTGACACGGAAC  
AACGGCAAACACGCCGCCGGGC

-----



Example of a many-to-one connectron in archea – *M. jannaschii*

In this example the existence of the T1-T2 (1630-1643) long loop is controlled by four C1/C2 short loops (1629, 1642, 124 and 1533).

5

1629 Chromosome 1  
1642 Chromosome 1  
124 Chromosome 1  
1533 Chromosome 1

10

\*-----\*-----\*  
|      Chromosome 1      |  
1630                              1643

15

A double stranded DNA loop of length 4.998 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 1630. This T1 control element has the DNA sequence

20

TTATTAATTAGTTCAAAGGATTTTTATTTAATTTCTAAGGGTTTGCTGGTTT  
GATTATTTAGAATATTTGAGTTTATTGAATTATTCAGATTTTAAAAATTA  
AGATTAATTAGGAAAGGAAATAAGATTTCTCTAACAGACAAGTTAAATTT  
TTGGATTTAAAAAGATAAAAAAT

25

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1643. This T2 control element has the DNA sequence

30

TTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTATTG  
AATTATTCAGATTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAAAT  
TTCTCTAACAAATAAGTTAAATTTTGGATTTAAAAAGATAAAAAATACTCT  
GTTTTATTATGGAAAGAAAGAT

35

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

MJ1597 MJ1598 MJ1599 MJ1600 MJ1601 MJ1602

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 1629 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1597 and has the DNA sequence

ATATGTTTGAAATTTGAAAATAAGAGTATTTAGAAGTTATTAATTAGTTCA  
AAGGATTTTTATTTAATTTCTAAGGGTTTGCTGGTTTGATTATTTAGAATAT  
TTGAGTTTATTGAATTATTCAGATTTTAAAAATTA

The match between the T1 sequence and the C1/C2 sequence is

TTATTAATTAGTTCAAAGGATTTTTATTTAATTTCTAAGGGTTTGCTGGTTT  
GATTATTTAGAATATTTGAGTTTATTGAATTATTCAGATTTTAAAAATTA

The match between the T2 sequence and the C1/C2 sequence is

GCTGGTTTGATTATTTAGAATATTTGAGTTTATTGAATTATTCAGATTTTAA  
AAAATTA

A C1/C2 short loop on chromosome 1 whose identifier is 1642 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1602 and has the DNA sequence

ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAA  
ATTTCTCTAACAAATAAGTTAAATTTTGGATTAAAAAGATAAAAATACT  
CTGTTTTATTATGGAAAGAAAGAT

The match between the T1 sequence and the C1/C2 sequence is

GCTGGTTTGATTATTTAGAATATTTGAGTTTATTGAATTATTCAGATTTTAA  
AAAATTA

The match between the T2 sequence and the C1/C2 sequence is

TTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTATTG  
AATTATTCAGATTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAAAT  
TTCTCTAACAAATAAGTTAAATTTTGGATTAAAAAGATAAAAATACTCT  
GTTTTATTATGGAAAGAAAGAT

A C1/C2 short loop on chromosome 1 whose identifier is 124 controls the expression  
of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA  
single strand that is 3'UTR to the gene MJ0112 and has the DNA sequence

ATTTAATTTCTAAGGGTTTGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTAAAAAT

The match between the T1 sequence and the C1/C2 sequence is

ATTTAATTTCTAAGGGTTTGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTAAAAAT

The match between the T2 sequence and the C1/C2 sequence is

GCTGGTTTGATTATTTAGAATATTTGAGTTTATTGAATTATTCAGATTTTAA  
AAAAT

A C1/C2 short loop on chromosome 1 whose identifier is 1533 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1486 and has the DNA sequence

5 TTTTATTTAATTTCTAAGGGTTTGCTGGTTTGATTATTTAGAATATTTGAG  
TTTATT

The match between the T1 sequence and the C1/C2 sequence is

10 TTTTATTTAATTTCTAAGGGTTTGCTGGTTTGATTATTTAGAATATTTGAG  
TTTATT

The match between the T2 sequence and the C1/C2 sequence is

15 GCTGGTTTGATTATTTAGAATATTTGAGTTTATT

-----

Example of a many-to-one connectron in single-cell eukaryotes – *S. cerevisiae*

20 In this example the existence of the T1-T2 (5515-5533) long loop on chromosome 12 is controlled by seventeen C1/C2 short loops (5516, 5532, 1939, 2323, 1942, 3286, 3649, 4764, 4751, 5536, 6102, 8023, 7356, 3293, 3291, 3289 and 146).

- 25 5516 Chromosome 12  
5532 Chromosome 12  
1939 Chromosome 4  
2323 Chromosome 5  
1942 Chromosome 5  
30 3286 Chromosome 7  
3649 Chromosome 8  
4764 Chromosome 12  
4751 Chromosome 12  
5536 Chromosome 13  
35 6102 Chromosome 14

5  
 8023 Chromosome 16  
 7356 Chromosome 16  
 3293 Chromosome 8  
 3291 Chromosome 8  
 3289 Chromosome 8  
 146 Chromosome 2  
 10  
 \*-----\*-----\*  
 | Chromosome 12 |  
 3197 3308

15  
 A double stranded DNA loop of length 6.466 kilo-bases on chromosome 12 is  
 bounded on the left by a T1 sequence whose identifier is 5515. This T1 control  
 element has the DNA sequence

20  
 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTTACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGTTG

25  
 This double stranded DNA loop is bounded on the right by a T2 control element  
 whose identifier is 5533. This T2 control element has the DNA sequence

30  
 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
 ATATGCGTTTTGATGTAGTAGTATTTACTGTTTTGATTTAGTGTTTGTGTC  
 ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
 TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGTTGAACA  
 TCCGGGTAAGAGACAACAGGGCT

35  
 This long T1/T2 double stranded DNA loop modulates the expression of the  
 following genes

YLR467W

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

5 A C1/C2 short loop on chromosome 12 whose identifier is 5516 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YLR464W and has the DNA sequence

10 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTAGTGTGTTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

15 A C1/C2 short loop on chromosome 12 whose identifier is 5532 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YLR467W and has the DNA sequence

20 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTAGTGTGTTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
25 AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

30 A C1/C2 short loop on chromosome 4 whose identifier is 1939 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed

as a RNA single strand that is 3'UTR to the gene YDR545W and has the DNA sequence

5 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGG

10 The match between the T1 sequence and the C1/C2 sequence is

15 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGG

The match between the T2 sequence and the C1/C2 sequence is

20 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTTCTAGGGA  
ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGG

25 A C1/C2 short loop on chromosome 5 whose identifier is 2323 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene YER189W and has the DNA  
sequence

30 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG

GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is

5

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTAGTGTGTTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
10 AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTTCTAGGGA  
15 ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTAGTGTGTTGTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
TCCGGGTA  
AGAGACAACAGGGCT

20

A C1/C2 short loop on chromosome 5 whose identifier is 1942 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene YEL077C and has the DNA  
sequence

25

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTAGTGTGTTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
30 AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is



AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 5 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

10 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTTCTAGGGA  
 ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTC  
 ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
 TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
 TCCGGGTA  
 15 AGAGACAACAGGGCT

A C1/C2 short loop on chromosome 7 whose identifier is 3286 controls the  
 expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
 as a RNA single strand that is 3'UTR to the gene YGR296W and has the DNA  
 20 sequence

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 25 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is

30 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG

GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

5

ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
TCCGGGTAAGAGACAACAGGGCT

10

A C1/C2 short loop on chromosome 8 whose identifier is 3649 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YHR219W and has the DNA sequence

15

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTTATTTAGTGTTTGTTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

20

The match between the T1 sequence and the C1/C2 sequence is

25

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTTATTTAGTGTTTGTTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTG

30

The match between the T2 sequence and the C1/C2 sequence is

ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
 ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTTC  
 ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
 TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
 5 TCCGGGTAAGAGACAACAGGGCT

A C1/C2 short loop on chromosome 12 whose identifier is 4764 controls the  
 expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
 as a RNA single strand that is 3'UTR to the gene YLL066C and has the DNA  
 10 sequence

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCCTGTTTTGATTTAGTGTTTGTTCACGGCAGTAGCGAGAGACAAGTG  
 15 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCCTGTTTTGATTTAGTGTTTGTTCACGGCAGTAGCGAGAGACAAGTG  
 20 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
 ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTTC  
 30 ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
 TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
 TCCGGGTAAGAGACAACAGGGCT

A C1/C2 short loop on chromosome 12 whose identifier is 4751 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YLL067C and has the DNA sequence

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTTGAATTTAGTGTGTTGTTGCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTTGAATTTAGTGTGTTGTTGCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTTCTAGGGA  
ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTGTTGTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
TCCGGGTAAGAGACAACAGGGCT

A C1/C2 short loop on chromosome 13 whose identifier is 5536 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YML133C and has the DNA sequence

5 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is

10 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

20 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTTTCTAGGGA  
 ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTC  
 ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
 TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
 TCCGGGTAAGAGACAACAGGGCT

25 A C1/C2 short loop on chromosome 14 whose identifier is 6102 controls the  
 expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
 as a RNA single strand that is 3'UTR to the gene YNL339C and has the DNA  
 sequence

30 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG

GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is

5

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTG

10

The match between the T2 sequence and the C1/C2 sequence is

ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
TCCGGGTAAGAGACAACAGGGCT

15

A C1/C2 short loop on chromosome 16 whose identifier is 8023 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YPR204W and has the DNA sequence

20

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

25

30

The match between the T1 sequence and the C1/C2 sequence is

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 5 AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

10 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTTCTAGGGA  
 ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTC  
 ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
 TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
 TCCGGGTAAGAGACAACAGGGCT

15 A C1/C2 short loop on chromosome 16 whose identifier is 7356 controls the  
 expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
 as a RNA single strand that is 3'UTR to the gene YPL283C and has the DNA  
 sequence

20 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

25

The match between the T1 sequence and the C1/C2 sequence is

30 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
 ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
 TTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
 GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
 AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

5 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
TCCGGGTAAGAGACAACAGGGCT

10 A C1/C2 short loop on chromosome 8 whose identifier is 3293 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene YHL050C and has the DNA  
sequence

15 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTT

The match between the T1 sequence and the C1/C2 sequence is

20 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTT

The match between the T2 sequence and the C1/C2 sequence is

25 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
ATATGCGTTTT

30 A C1/C2 short loop on chromosome 8 whose identifier is 3291 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene YHL050C and has the DNA  
sequence



ATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGC  
GAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAA

The match between the T1 sequence and the C1/C2 sequence is

5

ATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGC  
GAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAA

The match between the T2 sequence and the C1/C2 sequence is

10

ATGTAGTAGTATTTCACTGTTTTGATTTAGTGTTTGTGTCACGGCAGTAGC  
GAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAA

A C1/C2 short loop on chromosome 2 whose identifier is 145 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YBL113C and has the DNA sequence

15

CTATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAAC  
ATCCGGGTAAGAGACAACAGGCT

20

The match between the T1 sequence and the C1/C2 sequence is

CTATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTG

25

The match between the T2 sequence and the C1/C2 sequence is

CTATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAAC  
ATCCGGGTAAGAGACAACAGGCT

30

A C1/C2 short loop on chromosome 8 whose identifier is 3289 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed

as a RNA single strand that is 3'UTR to the gene YHL050C and has the DNA sequence

5 CTATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAAC  
ATCCGGGTAAGAGACAACAGGCT

The match between the T1 sequence and the C1/C2 sequence is

10 CTATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

15 CTATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAAC  
ATCCGGGTAAGAGACAACAGGCT

A C1/C2 short loop on chromosome 2 whose identifier is 146 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YBL113C and has the DNA sequence

20 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAA

The match between the T1 sequence and the C1/C2 sequence is

25 AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAA

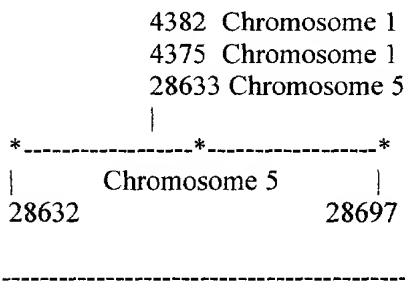
The match between the T2 sequence and the C1/C2 sequence is

30 ATTATGTATTGTGTAGTATAGTATATTGTAAGAAA

-----

Example of a many-to-one connectron in multi-cell eukaryotes – C. elegans

In this example the existence of the T1-T2 (3197-3308) long loop on chromosome 5 is controlled by three C1/C2 short loops (4382, 4375 and 28633).



A double stranded DNA loop of length 58.451 kilo-bases on chromosome 5 is bounded on the left by a T1 sequence whose identifier is 28632. This T1 control element has the DNA sequence

GCAAAAATTGACTGAAAATTTGAATTTCCCGCAAAAATTGACTGAAAAT  
TTGAATTTCCCGCCAAAATTGACTGAAAATTTGAA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 28697. This T2 control element has the DNA sequence

CAAAAATTGACTGAAAATTTGAATTTCCCTCCAAAATTGACTGAAAAT  
TTGAATTTCCCGCCAAAATTGACTGAAAATTTGAATATCCCGCCAAAAA  
TTGACTGAAAATTTGAATTTCCCGCCGAAAATTAAATGAAAATGGAATT  
TCTCGCCGAA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

M162.8 M162.4 M162.3 M162.6 M162.2 M162.1 M162.7

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

5

A C1/C2 short loop on chromosome 1 whose identifier is 4382 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene Y43F8B.10 and has the DNA sequence

10

ATTATAGAAAATTTAAATTTCCCTCCAAAAAATTGACTGAAAATTTGAATT  
TCCCTCCAAAAAATTGACTGAAAATTTGAATTTCCCGCCAAAAAATTGACTG  
AAAATTTGAATATCCCGCCAAAAAATTGACTGAAAATTTGAATTTCCCGCC  
GAAAATTAAATGAAAAATGGAATTTCTCGCCGAAAAATTCAGTAAAAATT  
15 TGAATTTCTTGCCAAAAAATTGACTGAAAATTTGAATTTCTTGCCAAAAA  
GTGACTGGGAATTTGAATTTCCCTCCAAAAAATTGACTGAAATTTTGAATTT  
CCCGCTAAAAGTTGACT

15

The match between the T1 sequence and the C1/C2 sequence is

20

CAAAAATTGACTGAAAATTTGAATTTCCCGC

The match between the T2 sequence and the C1/C2 sequence is

25

CAAAAATTGACTGAAAATTTGAATTTCCCTCCAAAAAATTGACTGAAAAT  
TTGAATTTCCCGCCAAAAAATTGACTGAAAATTTGAATATCCCGCCAAAAA  
TTGACTGAAAATTTGAATTTCCCGCCGAAAAATTAAATGAAAAATGGAATT  
TCTCGCCGAA

30

A C1/C2 short loop on chromosome 1 whose identifier is 4375 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed

as a RNA single strand that is 3'UTR to the gene Y43F8B.10 and has the DNA sequence

5 ATTATAGAAAATTTAAATTTCCCTCCAAAAAATTGACTGAAAATTTGAATT  
TCCCTCCAAAAATTGACTGAAAATTTGAATTTCCCGCCAAAAATTGACTG  
AAAATTTGAATATCCCGCCAAAAATTGACTGAAAATTTGAATTTCCCGCC  
GAAAATTAAATGAAAAATGGAATTTCTCGCCGAAAAATTCAGTAAAAATT  
TGAATTTCTTGCCAAAAATTGACTGAAAATTTGAATTTCTTGCCAAAAAA  
GTGACTGGGAATTTGAATTTCCCTCCAAAAATTGACTGAAATTTTGAATTT  
10 CCCGCTAAAAGTTGACT

The match between the T1 sequence and the C1/C2 sequence is

15 CAAAAATTGACTGAAAATTTGAATTTCCCGC

The match between the T2 sequence and the C1/C2 sequence is

20 CAAAAAATTGACTGAAAATTTGAATTTCCCTCCAAAAATTGACTGAAAAT  
TTGAATTTCCCGCCAAAAATTGACTGAAAATTTGAATATCCCGCCAAAAA  
TTGACTGAAAATTTGAATTTCCCGCCGAAAATTAAATGAAAAATGGAATT  
TCTCGCCGAA

25 A C1/C2 short loop on chromosome 5 whose identifier is 28633 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene M162.5 and has the DNA sequence

CAAAAAATTGACTGAAAATTTGAATTTCCCGCAAAAAATTGACTGAAAATT  
TGAATTTCCCGCCAAAAATTGACTGAAAATTTGAA

30 The match between the T1 sequence and the C1/C2 sequence is

CAAAAATTGACTGAAAATTTGAATTTCCCGCAAAAATTGACTGAAAATT  
TGAATTTCCCGCCAAAAATTGACTGAAAATTTGAA

The match between the T2 sequence and the C1/C2 sequence is

5

CAAAAATTGACTGAAAATTTGAATTTCCC

-----

10

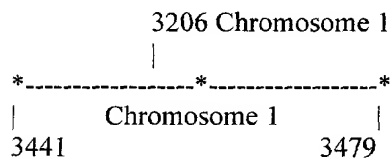
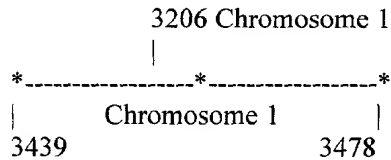
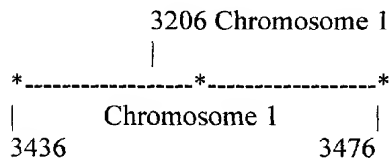
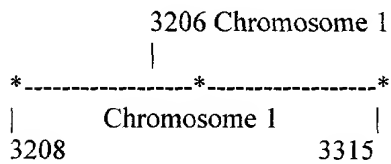
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3. One connectron controls the expression of many sets of genes in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes.

One C1/C2 short loop can control the existence of a many T1-T2 long loops. The C1/C2 short loop can be on the same chromosome or on different chromosomes from the T1-T2 long loops. This relationship is described as “one-to-many”. This relationship exists in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes.

Example of a one-to-many connectron in prokaryotes – E. coli

In this example the existence of T1-T2 (3208-3315, 3436-3476, 3439-3478 and 3441-3479) long loops are controlled by one C1/C2 short loop (3206).



-----

A double stranded DNA loop of length 93.377 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3208. This T1 control element has the DNA sequence

ACTCATCTTCGGGTGATGTTTGAGATATTTGCTCTTTAAAAATCTGGATCA  
 AGCTGAAAATTGAAACACTGAACAACGAAAGTTGTTTCGTGAGTCTCTCAA  
 ATTTTCGCAACACGATGATGAATCGAAAGAAACATCTTCGGGTTGTGAGG  
 TTAAGCGACTAAGCGTACACGGTGGATGCCCTGGC...AGTGTGTTTCGACA  
 CACTATCATTAAGTGAATCCATAGGTTAATGAGGCGAACCAGGGGGAACCTG  
 AAACATCTAAGTACCCCGAGGAAAAGAAATCAACCGAGATTCCCCCAGTA  
 GCGGCGAGCGAACGGGGAGCAGCCAGAGCCTGAATCAGT

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 3315. This T2 control element has the DNA sequence

TTTGCTCTTTAAAAATCTGGATCAAGCTGAAAATTGAAACACTGAACAAC  
 GAAAGTTGTTTCGTGAGTCTCTCAAATTTTCGCAACTCTGAAGTGAAACATC  
 TTCGGGTTGTGAGGTTAAGCGACTAAGCGTACACGGTGGATGCCCTGGCA  
 GTCAGAGGCGATGAAGGACGTGCTAATCTGCGATA...GGTTAATGAGGCG  
 AACCGGGGGAACCTGAAACATCTAAGTACCCCGAGGAAAAGAAATCAACC  
 GAGATTCCCCCAGTAGCGGCGAGCGAACGGGGAGCAGCCAGAGCCTGA  
 ATCAGTGTGTGTGTTAGTGGAAGCGTCTGGAAA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

rrlC	rrfC	aspT	trpT	yifA	yifE	yifB	ilvL	ilvG_1
ilvM	ilvE	ilvD	ilvA	ilvY	ilvC	ppiC	b3776	rep
gppA	rhlB	trxA	rhoL	rho	rfe	wzzE	wecB	rffH



wecD      wecE      wzxE      yifM\_2      wecG      yifK      argX      hisR  
 leuT      proM      aslB      aslA      hemY      hemX      hemD      cyaA  
 cyaY      b3808      dapF      uvrD      b3814      corA      yigF      yigG      rãrD  
 yigI      pldA      recQ      yigJ      yigK      pldB      yigL      yigM      metR  
 5      metE      ysgA      udp      yigN      ubiE      yigP      b3836      yigU  
 yigW\_1      rfaH      yigC      ubiB      fadA      fadB      pepQ      trkH  
 hemG      rrsA      ileT

10      The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

15      A C1/C2 short loop on chromosome 1 whose identifier is 3206 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene rrsC and has the DNA sequence

20      GTCCCCTTCGTCTAGAGGCCAGGACACCGCCCTTTCACGGCGGTAACAG  
 GGGTTCGAATCCCCTAGGGGACGCCACTTGCTGGTTTGTGAGTGAAAGTC  
 ACCTGCCTTAATATCTCAAACTCATCTTCGGGTGATGTTTGAGATATTTG  
 CTCTTTAAAAATCTGGATCAAGCTGAAAATTGAAA...ACCGGCGATTTCGG  
 AATGGGGAAACCCAGTGTGTTTCGACACACTATCATTAAGTGAATCCATA  
 GGTTAATGAGGCGAACCGGGGGAAGTGAACATCTAAGTACCCCGAGGA  
 AAAGAAATCAACCGAGATTCCCCCAGTAGCGGCGAGCGAACGGGGAGCA  
 GCCCAGAGCCTGAATCAGT

25      The match between the T1 sequence and the C1/C2 sequence is

30      ACTCATCTTCGGGTGATGTTTGAGATATTTGCTCTTTAAAAATCTGGATCA  
 AGCTGAAAATTGAAACACTGAACAACGAAAGTTGTTCGTGAGTCTCTCAA  
 ATTTTCGCAACACGATGATGAATCGAAAGAAACATCTTCGGGTTGTGAGG  
 TTAAGCGACTAAGCGTACACGGTGGATGCCCTGGC...AGTGTGTTTCGACA  
 CACTATCATTAAGTGAATCCATAGGTAAATGAGGCGAACCGGGGGAAGTGA

AAACATCTAAGTACCCCGAGGAAAAGAAATCAACCGAGATTCCCCCAGTA  
GCGGCGAGCGAACGGGGAGCAGCCCAGAGCCTGAATCAGT

The match between the T2 sequence and the C1/C2 sequence is

TTTGCTCTTTAAAAATCTGGATCAAGCTGAAAATTGAAACACTGAACAAC  
GAAAGTTGTTTCGTGAGTCTCTCAAATTTTCGCAAC

-----

A double stranded DNA loop of length 41.279 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3436. This T1 control element has the DNA sequence

ACGCAACGCGTGATAAGCAATTTTCGTGTCCCCTTCGTCTAGAGGCCAG  
GACACCGCCCTTTCACGGCGGTAACAGGGGTTCGAATCCCCTAGGGGACG  
CCTTGCTGGTT

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 3476. This T2 control element has the DNA sequence

AGTGAAGCAAGGCGTCTTGCGAAGCAGACTGATACGTCCCCTTCGTCT  
AGAGGCCAGGACACCGCCCTTTCACGGCGGTAACAGGGGTTCGAATCCC  
CTAGGGGACGCCACTTGCTGGTTTGTGAGTGAAAGTCACCTGCCTTAATA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

gltT	rrlB	rrfB	murB	coaA	b3975	tyrU	thrT	tufB
secE	nusG	rplK	rplA	rplJ	rplL	rpoB	rpoC	htrC
thiH	thiF	thiE	yjaE	yjaD	hemE	nfi	yjaG	hupA
yjaH	yjaI	hydH	purD	purH				

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

- 5 A C1/C2 short loop on chromosome 1 whose identifier is 3206 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene rrsC and has the DNA sequence

10 GTCCCCTTCGTCTAGAGGCCAGGACACCGCCCTTTCACGGCGGTAACAG  
GGGTTCTGAATCCCCTAGGGGACGCCACTTGCTGGTTTGTGAGTGAAAGTC  
ACCTGCCTTAATATCTCAAAACTCATCTTCGGGTGATGTTTGAGATATTTG  
CTCTTTAAAAATCTGGATCAAGCTGAAAATTGAAACACTGAACAACGAAA  
GTTGTTTCGTGAGTCTCTCAAATTTTCGCAACACGATGATGAATCGAAAGA  
AACATCTTCGGGTTGTGAGGTTAAGCGACTAAGCGTACACGGTGGATGCC  
15 CTGGCAGTCAGAGGCGATGAAGGACGTGCTAATCTGCGATAAGCGTCGGT  
AAGGTGATATGAACCGTTATAACCGGCGATTTCCGAATGGGGAAACCCAG  
TGTGTTTCGACACACTATCATTAAGTGAATCCATAGGTTAATGAGGCGAA  
CCGGGGGAACTGAAACATCTAAGTACCCCGAGGAAAAGAAATCAACCGA  
GATTCCCCCAGTAGCGGCGAGCGAACGGGGAGCAGCCCAGAGCCTGAAT  
20 CAGT

The match between the T1 sequence and the C1/C2 sequence is

25 GTCCCCTTCGTCTAGAGGCCAGGACACCGCCCTTTCACGGCGGTAACAG  
GGGTTCTGAATCCCCTAGGGGACGCCACTTGCTGGTT

The match between the T2 sequence and the C1/C2 sequence is

30 GTCCCCTTCGTCTAGAGGCCAGGACACCGCCCTTTCACGGCGGTAACAG  
GGGTTCTGAATCCCCTAGGGGACGCCACTTGCTGGTTTGTGAGTGAAAGTC  
ACCTGCCTTAATA

A double stranded DNA loop of length 41.336 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3439. This T1 control element has the DNA sequence

CCTTAATATCTCAAACTCATCTTCGGGTGATGTTTGAGATATTTGCTCTTT  
AAAAATCTGGATCAAGCTGAAAATTGAAACACTGAACAACGA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 3478. This T2 control element has the DNA sequence

GTGATGTTTGAGATATTTGCTCTTTAAAAATCTGGATCAAGCTGAAAATTG  
AAACACTGAACAACGAAAGTTGTTTCGTGAGTCTCTCAAATTTT

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

rrlB	rrfB	murB	coaA	b3975	tyrU	thrT	tufB	secE
nusG	rplK	rplA	rplJ	rplL	rpoB	rpoC	htrC	thiH
thiF	thiE	yjaE	yjaD	hemE	nfi	yjaG	hupA	yjaH
yjaI	hydH	purD	purH	gltV				

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 3206 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the generrC and has the DNA sequence

GTCCCCTTCGTCTAGAGGCCAGGACACCGCCCTTTCACGGCGGTAACAG  
GGGTTTGAATCCCCTAGGGGACGCCACTTGCTGGTTTGTGAGTGAAAGTC

ACCTGCCTTAATATCTCAAACTCATCTTCGGGTGATGTTTGAGATATTTG  
 CTCTTTAAAAATCTGGATCAAGCTGAAAATTGAAA...ACCGGCGATTTCGG  
 AATGGGGAAACCCAGTGTGTTTCGACACACTATCATTAAGTGAATCCATA  
 GGTTAATGAGGCGAACCGGGGGAAGTGAACATCTAAGTACCCCGAGGA  
 5 AAAGAAATCAACCGAGATTCCCCCAGTAGCGGCGAGCGAACGGGGAGCA  
 GCCCAGAGCCTGAATCAGT

The match between the T1 sequence and the C1/C2 sequence is

10 CCTTAATATCTCAAACTCATCTTCGGGTGATGTTTGAGATATTTGCTCTTT  
 AAAAATCTGGATCAAGCTGAAAATTGAAACACTGAACAACGA

The match between the T2 sequence and the C1/C2 sequence is

15 GTGATGTTTGAGATATTTGCTCTTTAAAAATCTGGATCAAGCTGAAAATTG  
 AAACACTGAACAACGAAAGTTGTTTCGTGAGTCTCTCAAATTTT

-----

20 A double stranded DNA loop of length 38.285 kilo-bases on chromosome 1 is  
 bounded on the left by a T1 sequence whose identifier is 3441. This T1 control  
 element has the DNA sequence

25 AATTTTCGCAACACGATGATGAATCGAAAGAAACATCTTCGGGTGTGAG  
 GTTAAGCGACTAAGCGTACACGGTGGATGCCCTGGCAGTCAGAGGCGATG  
 AAGGACGTGCTAATCTGCGATAAGCGTCGGTAAGGTGATATGAACCGTTA  
 TAACCGGCGATTTCCGAATGGGGAAACCCAGTGTGT...GATGAGAGAAGA  
 TTTTCAGCCTGATACAGATTAAATCAGAACGCAGAAGCGGTCTGATAAAA  
 CAGAATTTGCCTGGCGGCAGTAGCGCGGTGGTCCACCTGACCCCATGCC  
 30 GAACTCAGAAGTGAACGCCGTAGCGCCGATGGTAGTGTGGGGTCTCCCC  
 ATGCGAG

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 3479. This T2 control element has the DNA sequence

5 AAGAAACATCTTCGGGTTGTGAGGTTAAGCGACTAAGCGTACACGGTGGATGCCCTGGCAGTCAGAGGCGATGAAGGACGTGCTAATCTGCGATAAGCGTCGGTAAGGTGATATGAACCGTTATAACCGGCGATTTCCGAATGGGGAAACCCAGTGTGTTTCGACACACTATCATTAAGTGAATCC...CAGATTAAATCAGAACGCAGAAGCGGTCTGATAAAACAGAATTTGCCTGGCGGCAGTAGCGGGTGGTCCCACCTGACCCCATGCCGAAGTCAGAAGTGAAACGCCGTAGCGCCGATGGTAGTGTGGGGTCTCCCCATGCGAGAGTAGGGAAGTGCCAGGCA

10 TCAAATTA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

15

rrlB	rrfB	murB	coaA	b3975	tyrU	thrT	tufB	secE
nusG	rplK	rplA	rplJ	rplL	rpoB	rpoC	htrC	thiH
thiF	thiE	yjaE	yjaD	hemE	nfi	yjaG	hupA	yjaH
yjaI	hydH	purD	purH	gltV				

20 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

25 A C1/C2 short loop on chromosome 1 whose identifier is 3206 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene rrsC and has the DNA sequence

30 GTCCCCTTCGTCTAGAGGCCCAGGACACCGCCCTTTCACGGCGGTAACAGGGGTTTCGAATCCCCTAGGGGACGCCACTTGCTGGTTTGTGAGTGAAAGTCACTGCCTTAATATCTCAAACTCATCTTCGGGTGATGTTTGAGATATTTGCTCTTTAAAAATCTGGATCAAGCTGAAAATTGAAA...ACCGGCGATTTCCGAATGGGGAAACCCAGTGTGTTTCGACACACTATCATTAAGTGAATCCATA

GGTTAATGAGGCGAACCGGGGGAACCTGAAACATCTAAGTACCCCGAGGA  
AAAGAAATCAACCGAGATTCCCCCAGTAGCGGCGAGCGAACGGGGGAGCA  
GCCCAGAGCCTGAATCAGT

5 The match between the T1 sequence and the C1/C2 sequence is

AATTTTCGCAACACGATGATGAATCGAAAGAAACATCTTCGGGTTGTGAG  
GTTAAGCGACTAAGCGTACACGGTGGATGCCCTGGCAGTCAGAGGCGATG  
AAGGACGTGCTAATCTGCGATAAGCGTCGGTAAGGTGATATGAACCGTTA  
10 TAACCGGCGATTTCGAATGGGGAAACCCAGTGTGTTTCGACACACTATC  
ATTAAGTGAATCCATAGGTTAATGAGGCGAACCGGGGGAACCTGAAACATC  
TAAGTACCCCGAGGAAAAGAAATCAACCGAGATTCCCCCAGTAGCGGCG  
AGCGAACGGGGAGCAGCCCAGAGCCTGAATCAGT

15 The match between the T2 sequence and the C1/C2 sequence is

AAGAAACATCTTCGGGTTGTGAGGTTAAGCGACTAAGCGTACACGGTGG  
TGCCCTGGCAGTCAGAGGCGATGAAGGACGTGCTAATCTGCGATAAGCGT  
CGGTAAGGTGATATGAACCGTTATAACCGGCGATTTCGAATGGGGAAAC  
20 CCAGTGTGTTTCGACACACTATCATTAACTGAATCCATAGGTTAATGAGGC  
GAACCGGGGGAACCTGAAACATCTAAGTACCCCGAGGAAAAGAAATCAAC  
CGAGATTCCCCCAGTAGCGGCGAGCGAACGGGGAGCAGCCCAGAGCCTG  
AATCAGT

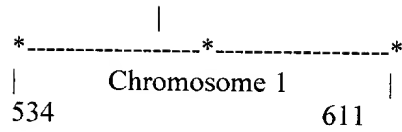
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Example of a one-to-many connectron in archea – *M. jannaschii*

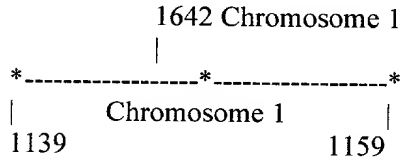
30 In this example the existence of T1-T2 (534-611, 1139-1159, and 1630-1643) long  
loops are controlled by one C1/C2 short loop (1642).

1642 Chromosome 1

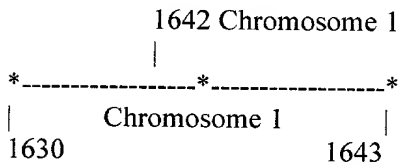
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15



20

A double stranded DNA loop of length 72.886 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 534. This T1 control element has the DNA sequence

25

TAAGTAAATAAAATTTCTCTAACAAATAAGTTAAATT

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 611. This T2 control element has the DNA sequence

30

TAAATAAAATTTCTCTAACAAATAAGTTAAATTTTGGATTAAAAAGATA  
AAAATGCT

35

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

MJ0486	MJ0487	MJ0488	MJ0489	MJ0490	MJ0492	MJ0493
MJ0494	MJ0495	MJ0496	MJ0497	MJ0499	MJ0500	MJ0501



MJ0502 MJ0503 MJ0504 MJ0506 MJ0507 MJ0508 MJ0509  
 MJ0510 MJ0511 MJ0512 MJ0513 MJ0514 MJ0514 MJ0517  
 MJ0519 MJ0520 MJ0521 MJ0522 MJ0523 MJ0525 MJ0526  
 MJ0526 MJ0529 MJ0530 MJ0531 MJ0532 MJ0534 MJ0535  
 5 MJ0536 MJ0538 MJ0539 MJ0540 MJ0541 MJ0542 MJ0543  
 MJ0544 MJ0545 MJ0547 MJ0548 MJ0549 MJ0550 MJ0552  
 MJ0553 MJ0554 MJ0555 MJ0556 MJ0558 MJ0559 MJ0560  
 MJ0561 MJ0562 MJ0563 MJ0564

10 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 1642 controls the  
 expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
 15 as a RNA single strand that is 3'UTR to the gene MJ1602 and has the DNA sequence

ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
 TGAATTATTCAGATTTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAA  
 ATTTCTCTAACAAATAAGTTAAATTTTTGGATTAAAAAAGATAAAAAATACT  
 20 CTGTTTTATTATGGAAAGAAAGAT

The match between the T1 sequence and the C1/C2 sequence is

25 AAGTAAATAAAATTTCTCTAACAAATAAGTTAAATT

The match between the T2 sequence and the C1/C2 sequence is

TAAATAAAATTTCTCTAACAAATAAGTTAAATTTTTGGATTAAAAAAGATA  
 AAAAT

30

-----

A double stranded DNA loop of length 14.509 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 1139. This T1 control element has the DNA sequence

5      ATTTATTAATTAGTTCAAAGGATTTTTATTTAATTTCTAAGGGTTAGCTGG  
TTTGATTGTTTAAAATATTTGAGTTTA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1159. This T2 control element has the DNA sequence

10      ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTTTAAAAATTA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

15      MJ1096    MJ1097    tRNA-Arg-3 MJ1098    MJ1099    MJ1100    MJ1101  
MJ1102    MJ1103    MJ1104    MJ1105    MJ1106    MJ1107    MJ1108

20      The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 1642 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1602 and has the DNA sequence

25      ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAA  
ATTTCTCTAACAAATAAGTTAAATTTTTGGATTAAAAAGATAAAAATACT  
30      CTGTTTTATTATGGAAAGAAAGAT

The match between the T1 sequence and the C1/C2 sequence is

ATTTAATTTCTAAGGGTTAGCTGGTTTGATT

The match between the T2 sequence and the C1/C2 sequence is

5

ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTAAAAATTA

-----

10

A double stranded DNA loop of length 4.998 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 1630. This T1 control element has the DNA sequence

15

TTATTAATTAGTTCAAAGGATTTTATTTAATTTCTAAGGGTTTGCTGGTTT  
GATTATTTAGAATATTTGAGTTTATTGAATTATTCAGATTTTAAAAATTA  
AGATTAATTAGGAAAGGAAATAAGATTTCTCTAACAGACAAGTTAAATTT  
TTGGATTTAAAAAGATAAAAAAT

20

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1643. This T2 control element has the DNA sequence

TTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTATTG  
AATTATTCAGATTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAAAT  
TTCTCTAACAAATAAGTTAAATTTTGGATTTAAAAAGATAAAAAATACTCT  
GTTTTATTATGGAAAGAAAGAT

25

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

30

MJ1597 MJ1598 MJ1599 MJ1600 MJ1601 MJ1602

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

5 A C1/C2 short loop on chromosome 1 whose identifier is 1642 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1602 and has the DNA sequence

10 ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAA  
ATTTCTCTAACAAATAAGTTAAATTTTTGGATTAAAAAGATAAAAAATACT  
CTGTTTTATTATGGAAAGAAAGAT

The match between the T1 sequence and the C1/C2 sequence is

15 GCTGGTTTGATTATTTAGAATATTTGAGTTTATTGAATTATTCAGATTTTTA  
AAAATTA

The match between the T2 sequence and the C1/C2 sequence is

20 TTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTATTG  
AATTATTCAGATTTTTTAAAAATTAGGATTAATTAGGCAAGTAAATAAAAT  
TTCTCTAACAAATAAGTTAAATTTTTGGATTAAAAAGATAAAAAATACTCT  
GTTTTATTATGGAAAGAAAGAT

25 -----

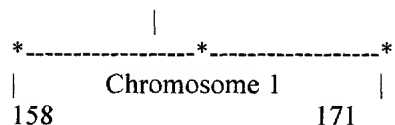
Example of a one-to-many connectron in single-cell eukaryotes – *S. cerevisiae*

30 In this example the existence of T1-T2 (158-171, 293-317, 4295-4308 and 5916-5923) long loops are controlled by one C1/C2 short loop (86).

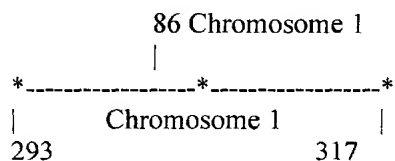
86 Chromosome 1

09866925.053001

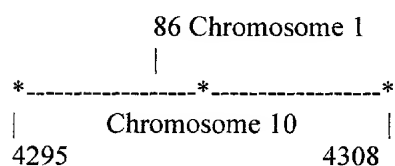
5



10

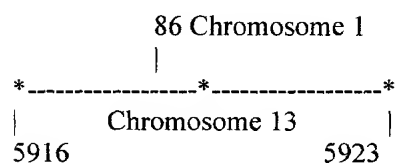


15



20

25



30

A double stranded DNA loop of length 20.391 kilo-bases on chromosome 2 is bounded on the left by a T1 sequence whose identifier is 158. This T1 control element has the DNA sequence

35

CCAATTGTTGGAATAAAAATCAACTATCATCTACTAACTAGTATTTACGTT  
ACTAGTATATTATCATATACGGTGTTAGAAGATGACGCAAATGATGAGAA  
ATAGTCATCTAAATTAGTGGAAGCTGAAACGCAAGGATTGATAATGTAAT  
AG

40

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 171. This T2 control element has the DNA sequence

ATAATTGTTGGAATAAAAATCAACTATCATCTACTAACTAGTATTTACGTT  
 ACTAGTATATTATCATATACGGTGTTAGAAGATGACACAAATGATGAGAA  
 ATAGTCATCTAAATTAGTGGAAGCTGAAACGCAAGGATTGATAATGTAAT  
 5 AGGATCAATGAATATTAACATATAAAAATGATGATAATAATA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

10 YBL107W-A TL(UAA)B1 YBL107C YBL106C YBL105C YBL104C  
 YBL103C YBL102W YBL101C

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

15 A C1/C2 short loop on chromosome 1 whose identifier is 86 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YAR009C and has the DNA sequence

20 ATCTATTACATTATGGGTGGTATGTTGGAATAGAAATCAACTATCATCTAC  
 TAACTAGTATTTACATTACTAGTATATTATCATATACGGTGTTAGAAGATG  
 ACGCAAATGATGAGAAATAGTCATCTAAATTAGTGGAAGCTGAAACGCA  
 AGGATTGATAATGTAATAGGATCAATGAATATAAACATATAAAACGGAAT  
 GAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGG  
 25 ATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTGTATACCTAATATT  
 ATAGCCTTTATCAACAATGGAATCCCAACAATTATCTCAACATTCACCCAT  
 TTCTCAGAA

The match between the T1 sequence and the C1/C2 sequence is

30 AAATCAACTATCATCTACTAACTAGTATTTAC

The match between the T2 sequence and the C1/C2 sequence is

AAATCAACTATCATCTACTAACTAGTATTTAC

5 -----

A double stranded DNA loop of length 38.470 kilo-bases on chromosome 2 is bounded on the left by a T1 sequence whose identifier is 293. This T1 control element has the DNA sequence

10

GAATTGTTGGAATAAAAATCCACTATCGTCTATCAACTAATAGTTATATTA  
TCAATATATTATCATATACGGTGTTAAGATGATGACATAAGTTATGAGAA  
GCTGTCATCGAAGTTAGAGGAAGCTGAAGTGCAAGGATTGATAATGTAAT  
AGGATAATGAAACATATAAAACGGAATGAGGAATAATCGTAATATTAGT  
15 ATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTTGAGGAGAAC  
TTCTAGT

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 317. This T2 control element has the DNA sequence

20

AATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTC  
GAGGAGAACTTCTAGTATATTCTGTA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

25

YBL005W-B TS(AGA)B YBL004W YBL003C YBL002W YBL001C  
YBR001C YBR002C YBR003W YBR004C YBR005W YBR006W  
YBR007C YBR008C YBR009C YBR010W YBR011C YBR012C

30

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 86 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YAR009C and has the DNA sequence

5

ATCTATTACATTATGGGTGGTATGTTGGAATAGAAATCAACTATCATCTAC  
TAACTAGTATTTACATTACTAGTATATTATCATATACGGTGTAGAAAGATG  
ACGCAAATGATGAGAAATAGTCATCTAAATTAGTGGAAGCTGAAACGCA  
AGGATTGATAATGTAATAGGATCAATGAATATAAACATATAAAACGGAAT  
10 GAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGG  
ATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTGTATACCTAATATT  
ATAGCCTTTATCAACAATGGAATCCCAACAATTATCTCAACATTCACCCAT  
TTCTCAGAA

15

The match between the T1 sequence and the C1/C2 sequence is

AAACATATAAAACGGAATGAGGAATAATCGTAATATTAGTATGTAGAAAT  
ATAGATTCCATTTTGAGGATTCCTATATCCT

20

The match between the T2 sequence and the C1/C2 sequence is

AATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTC  
GAGGAGAACTTCTAGTATATTCTGTA

25

-----

A double stranded DNA loop of length 11.020 kilo-bases on chromosome 10 is bounded on the left by a T1 sequence whose identifier is 4295. This T1 control element has the DNA sequence

30



AAACGCAAGGATTGATAATGTAATAGGATCAATGAATATAAACATATAAA  
ACGGAATGAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCAT  
TTTGAGGATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTG

- 5 This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 4308. This T2 control element has the DNA sequence

GGAAGCTGAAACGCAAGGATTGATAATGTAATAGGATCAATGAATATAA  
ACATATAAAACGGAATGAGGAATAATCGTAATATTAGTATGTAGAAATAT  
10 AGATTCCATTTTGAGGATTCCTATATCCTCGAGGAGAACTTCTAGTATATT  
CTGTATACCTAATATTATAGCCTTTATCAA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

15 YJR027W YJR029W

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

20 A C1/C2 short loop on chromosome 1 whose identifier is 87 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YAR009C and has the DNA sequence

25 ATCTATTACATTATGGGTGGTATGTTGGAATAGAAATCAACTATCATCTAC  
TAACTAGTATTTACATTACTAGTATATTATCATATACGGTGTTAGAAGATG  
ACGCAAATGATGAGAAATAGTCATCTAAATTAGTGGAAGCTGAAACGCA  
AGGATTGATAATGTAATAGGATCAATGAATATAAACATATAAAACGGAAT  
GAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGG  
30 ATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTGTATACCTAATATT  
ATAGCCTTTATCAACAATGGAATCCCAACAATTATCTCAACATTCACCCAT  
TTCTCA

-----

A double stranded DNA loop of length 5.462 kilo-bases on chromosome 13 is bounded on the left by a T1 sequence whose identifier is 5916. This T1 control element has the DNA sequence

AAGCTGAAGTGCAAGGATTGATAATGTAATAGGATAATGAAACATATAA  
AACGGAATGAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCA  
TTTTGAGGATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTGTA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 5923. This T2 control element has the DNA sequence

TAATAGGATAATGAAACATATAAAACGGAATGAGGAATAATCGTAATATT  
AGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTCGAGGAG  
AACTTCTAGTATATTCTGTATACCTAATATTATAGCCTTTATCAA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

YML045W

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 87 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YAR009C and has the DNA sequence

ATCTATTACATTATGGGTGGTATGTTGGAATAGAAATCAACTATCATCTAC  
TAACTAGTATTTACATTACTAGTATATTATCATATACGGTGTTAGAAGATG

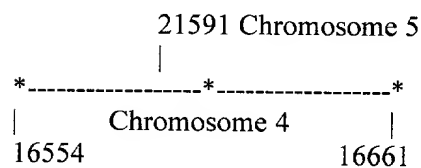
ACGCAAATGATGAGAAATAGTCATCTAAATTAGTGGAAGCTGAAACGCA  
 AGGATTGATAATGTAATAGGATCAATGAATATAAACATATAAAACGGAAT  
 GAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGG  
 ATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTGTATACCTAATATT  
 5 ATAGCCTTTATCAACAATGGAATCCCAACAATTATCTCAACATTCACCCAT  
 TTCTCA

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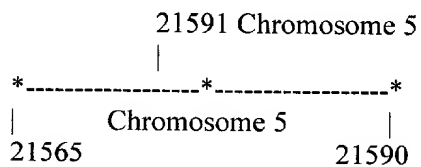
10 Example of a one-to-many connectron in multi-cell eukaryotes – *C. elegans*

In this example the existence of T1-T2 (16554-16661 and 21565-21590) long loops  
 are controlled by one C1/C2 short loop (21591).

15



25



30

-----

35 A double stranded DNA loop of length 50.159 kilo-bases on chromosome 4 is  
 bounded on the left by a T1 sequence whose identifier is 16554. This T1 control  
 element has the DNA sequence

TGCCTGAAAAAATTGGCTCCGAGTTAGGACACTTGGGGTGGTCAAAAAAT  
TTTGTGACTATTGTCAAATGAAAGATCATAGTTGATAACATAAATTCCCAA  
AGTTTCATAAAAATCGATACGCAGCGAACAAGTTATCAATT

- 5 This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 16661. This T2 control element has the DNA sequence

CACTTGGGGTGGTCAAAAAATTTTGTGATTATTGTCAAATGAAAGATCAT  
GGTTGATAACATAAATTCCCAAAGTTTCATAAAAATCGATACGCAGCGAA  
10 CAAAGTTATGATTTTTGACCCGGAAGTTATTTGGAGACCTA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

- 15 C23H5.7 C23H5.8a C23H5.3 C23H5.2 C23H5.9 C23H5.1

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

- 20 A C1/C2 short loop on chromosome 5 whose identifier is 21591 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene F25A2.1 and has the DNA sequence

TATTGTCAAATGAAAGATCATGGTTGATAACATAAATTCACAAATTTTCAT  
25 AAAAATCGATACGCAGCGAACAAGTTATGATTTTTGACCCGGAAGTTAT  
TTGGAGACCTAATATT

The match between the T1 sequence and the C1/C2 sequence is

- 30 TTTCATAAAAATCGATACGCAGCGAACAAGTTAT

The match between the T2 sequence and the C1/C2 sequence is

TATTGTCAAATGAAAGATCATGGTTGATAACATAAAATTCCCA

5 -----

A double stranded DNA loop of length 18.142 kilo-bases on chromosome 5 is bounded on the left by a T1 sequence whose identifier is 21565. This T1 control element has the DNA sequence

10

CTCCGAGTTAGGACACTTGGGGTGGACAAAAAATTTTGTGACTATTGTCA  
AATGAAAGATCATGGTTGATAA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 21590. This T2 control element has the DNA sequence

15

TATTGTCAAATGAAAGATCATGGTTGATAACATAAAATTTCCACAATTTTCAT  
AAAAATCGATACGCAGCGAACAAAGTTATGATTTTTGACCCGGAAGTTAT  
TTGGAGACCTAATA

20

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

T21H3.2 T21H3.1 F25A2.1

25

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 5 whose identifier is 21591 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene F25A2.1 and has the DNA sequence

30

TATTGTCAAATGAAAGATCATGGTTGATAACATAAATTCCCACAATTTTCAT  
AAAAATCGATACGCAGCGAACAAAGTTATGATTTTTGACCCGGAAGTTAT  
TTGGAGACCTAATATT

5 The match between the T1 sequence and the C1/C2 sequence is

TATTGTCAAATGAAAGATCATGGTTGATAA

The match between the T2 sequence and the C1/C2 sequence is

10

TATTGTCAAATGAAAGATCATGGTTGATAACATAAATTCCCACAATTTTCAT  
AAAAATCGATACGCAGCGAACAAAGTTATGATTTTTGACCCGGAAGTTAT  
TTGGAGACCTAATA

15

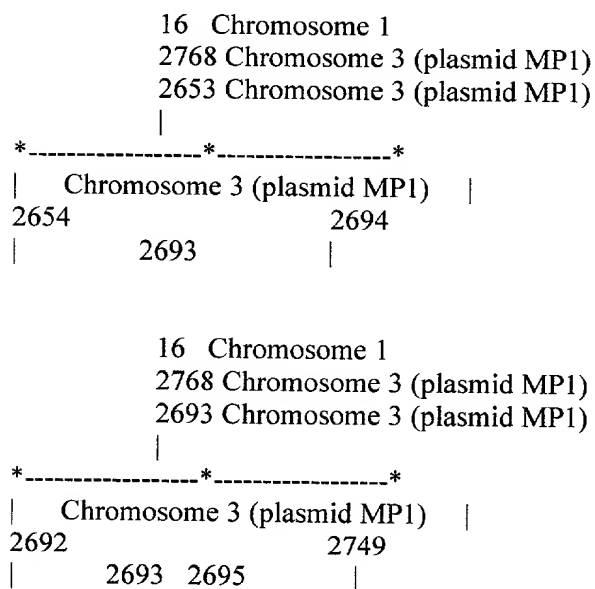
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4. Connectrons occur between prokaryotes and their plasmids.

Connectron relationships exist between prokaryotes and their plasmids. These connectrons implement a control mechanism between the two genomes that makes it possible for them to form a symbiotic relationship. In the case of *D. radiodurans* the relationship is not symmetric. The *D. radiodurans* genome sends C1/C2 short loops to the MP1 plasmid.

Example of a prokaryote/plasmid connectron – *D. radiodurans*

In this example the existence of T1-T2 (2654-2694 and 2692-2749) long loops in chromosome 3 that is the plasmid MP1 are controlled by one C1/C2 short loop (16) in chromosome 1.



A double stranded DNA loop of length 46.903 kilo-bases on chromosome 3 (plasmid MP1) is bounded on the left by a T1 sequence whose identifier is 2654. This T1 control element has the DNA sequence

CAGCGTTTTCTCGCTGTTCTGACGGCTGAACGCCCTGAATCTCTCCCG  
 GTATGCAGCCTGCTCGGAGAGTACGATTCGTCGTTGGCTGCACCGAAGTG  
 ACGATGGGGCCATTCCGTGGGGCGCGTTACACCAGGCGACTGTCAGTACA  
 5 GCAATCGAGAGTGGGCTGATCAGCCCACTGTGCGTTCTGGCCATCGACGC  
 CTCTTTTCACCGCAAAGCCGGTCAGCACACCGCACACCTCGGCTCGTTCTG  
 GAATGGCTGTGCCGCGCGGACC

10 This double stranded DNA loop is bounded on the right by a T2 control element  
 whose identifier is 2694. This T2 control element has the DNA sequence

GCTGAACGCCCTGAATCTCTCCCGGTATGCAGCCTGCTCGGAGAGTACGA  
 TTCGTCGTTGGCTGCACCGAAGTGACGATGGGGCCATTCCGTGGGGCGCG  
 TTACACCAGGCGACTGTCAGTACAGCAATCGAGAGTGGGCTGATCAGCCC  
 15 ACTGTGCGTTCTGGCCATCGACGCCTCTTTTCACCGCAAAGCCGGTCAGCA  
 CACCGCACACCTCGGCTCGTTCTGGAATGGCTGTGCCGCGCGGACCGAAC  
 GCGGAATCGAGCAATCCTGTTGT

20 This long T1/T2 double stranded DNA loop modulates the expression of the  
 following genes

DRB0020	DRB0021	DRB0022	DRB0023	DRB0024	DRB0025
DRB0027	DRB0030	DRB0032	DRB0033	DRB0034	DRB0035
DRB0037	DRB0038	DRB0039	DRB0041	DRB0042	DRB0043
25 DRB0044	DRB0045	DRB0047	DRB0051	DRB0052	DRB0054
DRB0055	DRB0057				

30 This long T1/T2 double stranded DNA loop modulates the expression of the  
 following C1/C2 short loops

A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2693  
 controls the expression of the genes of one or more other T1/T2 long loops. This



C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DRB0057 and has the DNA sequence

5 CTGATGGCCATCCTACAGTACGTTCTCAGCGCGGTCCCGCTGCGCAAGAC  
GCAGCGGAATTCCTGACCGTGCTGCTCAGCGTTTTTCTCGCTGTTCTCTGG  
AC

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

10

A C1/C2 short loop on chromosome 1 whose identifier is 16 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DR0009 and has the DNA sequence

15 GCTGTGAAATCACCGCTTCCAATGGGTCTGATGGCCATCCTACAGTACGTT  
CTCAGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCT  
GCTCAGCGTTTTTCTCGCTGTTCTCTGGACGGCTGAACGCCCTGAATCTCTC  
CCGGTATGCAGCCTGCTCGGAGAGTACGATTCGT

20

The match between the T1 sequence and the C1/C2 sequence is

25 CAGCGTTTTTCTCGCTGTTCTCTGGACGGCTGAACGCCCTGAATCTCTCCCG  
GTATGCAGCCTGCTCGGAGAGTACGATTCGTTCGTTGGCTGCACCGAAGTG  
ACGATGGGGCCATTCCGTGGGGCGCGTTACACCAGGCGACTGTCAGTACA  
GCAATCGAGAGTGGGCTGATCAGCCCACTGTGCGTTCTGGCCATCGACGC  
CTCTTTTCACCGCAAAGCCGGTCAGCACACCGCACACCTCGGCTCGTTCTG  
GAATGGCTGTGCCGCGCGGACC

30

The match between the T2 sequence and the C1/C2 sequence is

GCTGAACGCCCTGAATCTCTCCCGGTATGCAGCCTGCTCGGAGAGTACGA  
 TTCGTCGTTGGCTGCACCGAAGTGACGATGGGGCCATTCCGTGGGGCGCG  
 TTACACCAGGCGACTGTCAGTACAGCAATCGAGAGTGGGCTGATCAGCCC  
 ACTGTGCGTTCTGGCCATCGACGCCTCTTTTCACCGCAAAGCCGGTCAGCA  
 5 CACCGCACACCTCGGCTCGTTCTGGAATGGCTGTGCCGCGCGGACCGAAC  
 GCGGAATCGAGCAATCCTGTTGT

A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2768  
 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is  
 10 expressed as a RNA single strand that is 3'UTR to the gene DRB0133 and has the  
 DNA sequence

GCTGTGAAATCACCGCTTCCAATGGGTCTGATGGCCATCCTACAGTACGTT  
 CTCAGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTTCTGACCGTGCT  
 15 GCTCAGCGTTTTTCTCGCTGTTCTGGACGGCTGAACGCCCTGAATCTCTC  
 CCGGTATGCAGCCTGCTCGGAGAGTACGATTCGT

The match between the T1 sequence and the C1/C2 sequence is

20 CAGCGTTTTTCTCGCTGTTCTGGACGGCTGAACGCCCTGAATCTCTCCCG  
 GTATGCAGCCTGCTCGGAGAGTACGATTCGTGCTGTTGGCTGCACCGAAGTG  
 ACGATGGGGCCATTCCGTGGGGCGCGTTACACCAGGCGACTGTCAGTACA  
 GCAATCGAGAGTGGGCTGATCAGCCCACTGTGCGTTCTGGCCATCGACGC  
 25 CTCTTTTCACCGCAAAGCCGGTCAGCACACCGCACACCTCGGCTCGTTCTG  
 GAATGGCTGTGCCGCGCGGACC

The match between the T2 sequence and the C1/C2 sequence is

30 GCTGAACGCCCTGAATCTCTCCCGGTATGCAGCCTGCTCGGAGAGTACGA  
 TTCGTCGTTGGCTGCACCGAAGTGACGATGGGGCCATTCCGTGGGGCGCG  
 TTACACCAGGCGACTGTCAGTACAGCAATCGAGAGTGGGCTGATCAGCCC

ACTGTGCGTTCTGGCCATCGACGCCTCTTTTCACCGCAAAGCCGGTCAGCA  
 CACCGCACACCTCGGCTCGTTCTGGAATGGCTGTGCCGCGCGGACCGAAC  
 GCGGAATCGAGCAATCCTGTTGT

5 A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2653 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DRB0017 and has the DNA sequence

10 CGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCTGCTCAGC  
 GTTTTCTCGCTGTTCTGGACGGCTGAACGCCCTGAATCTCTCCCGGTAT  
 GCAGCCTGCTCGGAGAGTACGATTCGTCGTTGGCTGCACCGAAGTGACGA  
 TGGGGCCATTCCGTGGGGCGCGTTACACCAGGCGA

15 The match between the T1 sequence and the C1/C2 sequence is

CAGCGTTTTTCTCGCTGTTCTGGACGGCTGAACGCCCTGAATCTCTCCCG  
 GTATGCAGCCTGCTCGGAGAGTACGATTCGTCGTTGGCTGCACCGAAGTG  
 ACGATGGGGCCATTCCGTGGGGCGCGTTACACCAGGCGACTGTCAGTACA  
 20 GCAATCGAGAGTGGGCTGATCAGCCCACTGTGCGTTCTGGCCATCGACGC  
 CTCTTTTCACCGCAAAGCCGGTCAGCACACCGCACACCTCGGCTCGTTCTG  
 GAATGGCTGTGCCGCGCGGACC

The match between the T2 sequence and the C1/C2 sequence is

25 GCTGAACGCCCTGAATCTCTCCCGGTATGCAGCCTGCTCGGAGAGTACGA  
 TTCGTCGTTGGCTGCACCGAAGTGACGATGGGGCCATTCCGTGGGGCGCG  
 TTACACCAGGCGACTGTCAGTACAGCAATCGAGAGTGGGCTGATCAGCCC  
 ACTGTGCGTTCTGGCCATCGACGCCTCTTTTCACCGCAAAGCCGGTCAGCA  
 30 CACCGCACACCTCGGCTCGTTCTGGAATGGCTGTGCCGCGCGGACCGAAC  
 GCGGAATCGAGCAATCCTGTTGT

-----

A double stranded DNA loop of length 68.612 kilo-bases on chromosome 3 (plasmid MP1) is bounded on the left by a T1 sequence whose identifier is 2692. This T1 control element has the DNA sequence

CTGATGGCCATCCTACAGTACGTTCTCAGCGCGGTCCCGCTGCGCAAGAC  
GCAGCGGAATTCCTGACCGTGCTGCTCAGCGTTTTTCTCGCTGTTCCCTGG  
AC

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 2749. This T2 control element has the DNA sequence

AGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCTGCT  
CAGCGTTTTTCTCGCTGTTCCCTGGACGGCTGAACGCCCTGAATCTCTCCCG  
GT

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

DRB0059	DRB0060	DRB0061	DRB0062	DRB0064	DRB0065
DRB0066	DRB0067	DRB0068	DRB0069	DRB0070	DRB0072
DRB0073	DRB0074	DRB0076	DRB0077	DRB0079	DRB0080
DRB0081	DRB0083	DRB0085	DRB0086	DRB0087	DRB0088
DRB0089	DRB0090	DRB0092	DRB0093	DRB0094	DRB0096
DRB0097	DRB0098	DRB0102	DRB0103	DRB0104	DRB0105
DRB0106	DRB0107	DRB0111	DRB0112		

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2693 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DRB0057 and has the DNA sequence

5

CTGATGGCCATCCTACAGTACGTTCTCAGCGCGGTCCCGCTGCGCAAGAC  
GCAGCGGAATTTTCCTGACCGTGCTGCTCAGCGTTTTTCTCGCTGTTCTCTGG  
AC

10

A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2695 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DRB0057 and has the DNA sequence

15

GCTGAACGCCCTGAATCTCTCCCGGTATGCAGCCTGCTCGGAGAGTACGA  
TTCGTCGTTGGCTGCACCGAAGTGACGATGGGGCCATTCCGTGGGGCGCG  
TTACACCAGGCGACTGTCAGTACAGCAATCGAGAGTGGGCTGATCAGCCC  
ACTGTGCGTTCTGGCCATCGACGCCTCTTTTCACCGCAAAGCCGGTCAGCA  
CACCGCACACCTCGGCTCGTTCTGGAATGGCTGTGCCGCGCGGACCGAAC  
GCGGAATCGAGCAATCCTGTTGT

20

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

25

A C1/C2 short loop on chromosome 1 whose identifier is 16 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DR0009 and has the DNA sequence

30

GCTGTGAAATCACCGCTTCCAATGGGTCTGATGGCCATCCTACAGTACGTT  
CTCAGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTTTCCTGACCGTGCT  
GCTCAGCGTTTTTCTCGCTGTTCTCTGGACGGCTGAACGCCCTGAATCTCTC  
CCGGTATGCAGCCTGCTCGGAGAGTACGATTTCGT

The match between the T1 sequence and the C1/C2 sequence is

5 CTGATGGCCATCCTACAGTACGTTCTCAGCGCGGTCCCGCTGCGCAAGAC  
GCAGCGGAATTTCTGACCGTGCTGCTCAGCGTTTTTCTCGCTGTTCTCTGG  
AC

The match between the T2 sequence and the C1/C2 sequence is

10 AGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTTCTGACCGTGCTGCT  
CAGCGTTTTTCTCGCTGTTCTGGACGGCTGAACGCCCTGAATCTCTCCCG  
GT

15 A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2768  
controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is  
expressed as a RNA single strand that is 3'UTR to the gene DRB0133 and has the  
DNA sequence

20 GCTGTGAAATCACCGCTTCCAATGGGTCTGATGGCCATCCTACAGTACGTT  
CTCAGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTTCTGACCGTGCT  
GCTCAGCGTTTTTCTCGCTGTTCTGGACGGCTGAACGCCCTGAATCTCTC  
CCGGTATGCAGCCTGCTCGGAGAGTACGATTCTGT...CGGACCGAACGCGGA  
ATCGAGCAATCCTGTTGTGCCCTCATTGATGTCCAGCACCGGCAGGCCTTG  
ACGGTCGATGTCCGTCAGACCCTGACCGGGTCTGAGGCTCCAACTCGTCT  
25 GGAACAG

The match between the T1 sequence and the C1/C2 sequence is

30 CTGATGGCCATCCTACAGTACGTTCTCAGCGCGGTCCCGCTGCGCAAGAC  
GCAGCGGAATTTCTGACCGTGCTGCTCAGCGTTTTTCTCGCTGTTCTCTGG  
AC

The match between the T2 sequence and the C1/C2 sequence is

AGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCTGCT  
CAGCGTTTTTCTCGCTGTTCTGACGGCTGAACGCCCTGAATCTCTCCCG  
GT

A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2693 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DRB0057 and has the DNA sequence

CTGATGGCCATCCTACAGTACGTTCTCAGCGCGGTCCCGCTGCGCAAGAC  
GCAGCGGAATTCCTGACCGTGCTGCTCAGCGTTTTTCTCGCTGTTCTGGA  
AC

The match between the T1 sequence and the C1/C2 sequence is

CTGATGGCCATCCTACAGTACGTTCTCAGCGCGGTCCCGCTGCGCAAGAC  
GCAGCGGAATTCCTGACCGTGCTGCTCAGCGTTTTTCTCGCTGTTCTGGA  
AC

The match between the T2 sequence and the C1/C2 sequence is

AGCGCGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCTGCT  
CAGCGTTTTTCTCGCTGTTCTGGA

A C1/C2 short loop on chromosome 3 (plasmid MP1) whose identifier is 2653 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene DRB0017 and has the DNA sequence

CGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCTGCTCAGC  
GTTTTTCTCGCTGTTTCCTGGACGGCTGAACGCCCTGAATCTCTCCCGGTAT  
GCAGCCTGCTCGGAGAGTACGATTCGTCGTTGGCTGCACCGAAGTGACGA  
TGGGGCCATTCCGTGGGGCGCGTTACACCAGGCGA

5

The match between the T1 sequence and the C1/C2 sequence is

CGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCTGCTCAGC  
GTTTTTCTCGCTGTTTCCTGGAC

10

The match between the T2 sequence and the C1/C2 sequence is

CGGTCCCGCTGCGCAAGACGCAGCGGAATTCCTGACCGTGCTGCTCAGC  
GTTTTTCTCGCTGTTTCCTGGACGGCTGAACGCCCTGAATCTCTCCCGGT

15

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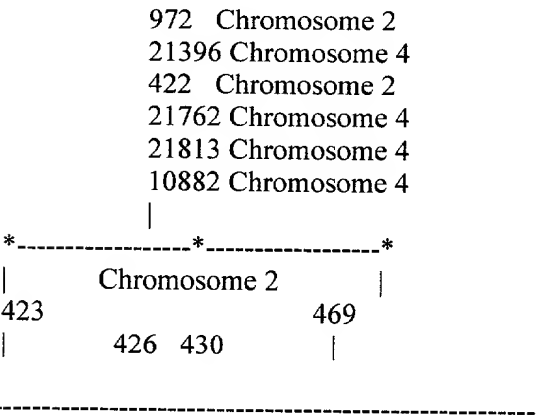


5. Connectrons occur in plants and higher animals

Connectron relationships exist in plant and higher animals.

5 Example of a plant connectron – *A. thaliana*

In this example the existence of the T1-T2 (423-469) long loop is controlled by six C1/C2 short loops (972, 21396, 422, 21762, 21813 and 10882). The T1-T2 long loop controls the expression of six genes on chromosome 2 in addition to two C1/C2 (426 and 430) short loops.



A double stranded DNA loop of length 42.285 kilo-bases on chromosome 2 is bounded on the left by a T1 sequence whose identifier is 423. This T1 control element has the DNA sequence

TATCTCTTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTA  
ATTAAAAAACGAAATA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 469. This T2 control element has the DNA sequence

TACTAATTTAATTAATTAATTAATTTAATTAACGAAATACATTATTAATT  
TTCAAAAATAATAACC

5 This long T1/T2 double stranded DNA loop modulates the expression of the following genes

At2g02070 At2g02080 At2g02090 At2g02100 At2g02120 At2g02130

10 This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

15 A C1/C2 short loop on chromosome 2 whose identifier is 426 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene At2g02060 and has the DNA sequence

TTCCAAAATAATAACCAATCAAAATCAACATATAAGATTTGATATCTAA  
ATTTT

20 A C1/C2 short loop on chromosome 2 whose identifier is 430 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene At2g02060 and has the DNA sequence

25 TTGCGGAAAAATAATATCATCATTATAAAAAATAATTAGAGTTTTTCGC  
ATAT

30 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 2 whose identifier is 972 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene At2g04240 and has the DNA sequence

5 GTATGCCATTAGAAATAAAATTTTAAAAGTAAATTAATTCATCTCTTTAAA  
AATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTAATTAAAAAACG  
AAATACATTATTAATTT

The match between the T1 sequence and the C1/C2 sequence is

10 ATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTAATTAAAAAACGA  
AATA

The match between the T2 sequence and the C1/C2 sequence is

15 TACTAATTTAATTAATTAAATTTAATTAAAAAACGAAATACATTATTAATT  
T

A C1/C2 short loop on chromosome 4 whose identifier is 21396 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene AT4g15300 and has the DNA sequence

20 TGCCATTAGAAATAAAATTTTAAAGAGTAAATTAATTTATCTCTTTAAGGA  
25 TTAAAAAGTCAAATACTAATTTAATTAATTAAATTTAATTAAAAAACGAA  
ATACATTATTAATTTCCAAAA

The match between the T1 sequence and the C1/C2 sequence is

30 TATCTCTTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTA  
ATTAAAAAACGAAATA

The match between the T2 sequence and the C1/C2 sequence is

TACTAATTTAATTAATTAAATTTAATTAAAAACGAAATACATTATTAATT  
T

5

A C1/C2 short loop on chromosome 2 whose identifier is 422 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene At2g02060 and has the DNA sequence

10

TAACCTTAATTTTTGTAAGTAATTATATAGGTATGCCATTAGAAATAAAAT  
TTTAAAGAGTAAATTAATTTATCTCTTTAAGGATTAAAAAGTCAAATACTA  
ATTTAATTAATTAAATTTAATTAAAAACGAAATA

The match between the T1 sequence and the C1/C2 sequence is

15

TATCTCTTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTA  
ATTAAAAACGAAATA

The match between the T2 sequence and the C1/C2 sequence is

20

TACTAATTTAATTAATTAAATTTAATTAAAAACGAAATA

A C1/C2 short loop on chromosome 4 whose identifier is 21762 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene AT4g17510 and has the DNA sequence

25

TTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTAATTAAA  
AACGAAATACATT

30

The match between the T1 sequence and the C1/C2 sequence is

TTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTAATTAAA  
AAACGAAATA

The match between the T2 sequence and the C1/C2 sequence is

TACTAATTTAATTAATTAAATTTAATTAAAAAACGAAATACATT

A C1/C2 short loop on chromosome 4 whose identifier is 21813 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene AT4g17680 and has the DNA sequence

TTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTAATTAAA  
AAACGAAATACATT

The match between the T1 sequence and the C1/C2 sequence is

TTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTAATTAAA  
AAACGAAATA

The match between the T2 sequence and the C1/C2 sequence is

TACTAATTTAATTAATTAAATTTAATTAAAAAACGAAATACATT

A C1/C2 short loop on chromosome 2 whose identifier is 10882 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene At2g26540 and has the DNA sequence

TATCTCTTTAAGGATTAAAAAGTCAAATACTAATTTAATTAATTAAATTTA  
ATTAA



- - - - -  
 - CG11207 - CG2186 CG2157  
 - Ork1 - - -  
 -

5

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

10 A C1/C2 short loop on chromosome 4 whose identifier is 3362 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene XXX and has the DNA sequence

15 AAAAAAGTACCGCGTTTTACTCCTAATTACCAATTCTAACCATCCATATCA  
 CTTTTTGACGGACTCCGTTAAAATAATTTTTGACCAAATTTTCGCATTTTTT  
 GTAATCAAAATTTGCAAAAAATTGAAAAAAC

20 A C1/C2 short loop on chromosome 4 whose identifier is 3364 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene XXX and has the DNA sequence

25 CAAAATTTGAATGCAAATCGATTGGGAATCAAAAAACAAACTCAACGAG  
 GTATGACATTCCATATTTGGGCCATTATTTCCAA

30 A C1/C2 short loop on chromosome 4 whose identifier is 3366 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene XXX and has the DNA sequence

TTTTTTCACAAAAATTAGGAAAATGATTTTGGGTAAAAAAATGAATATTT  
AAGTTGGGTTTT

5 A C1/C2 short loop on chromosome 4 whose identifier is 3369 controls the  
expression of the genes of one or more other T1/T2 long loops. This C1/C2 short  
loop is expressed as a RNA single strand that is 3'UTR to the gene XXX and has the  
DNA sequence

10 AAATCGATTGGGAATCAAAAAACAAACCTCAACGAGGTATGACATTCCAT  
ATCTGGGCCATTATTTCCAATCTTTTGATCAAAATAC

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2  
short loops.

15 A C1/C2 short loop on chromosome 4 whose identifier is 3373 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene XXX and has the DNA sequence

20 AAAAAAGTACCGCGTTTTACTCCTAATTACCAATTCTAACCATCCATATCA  
CTTTTGTACGGACTCCGTGAAAATAATTTTGGCCAAATTTTCGCATTTTTT  
GTAAGGGGTAACATCATCAAAATTTGCGAAAAA

The match between the T1 sequence and the C1/C2 sequence is

25 TTTTACTCCTAATTACCAATTCTAACCATCCATATCACTTTTTGACGGACTC  
CGTGAAAATAATTTTGGCCAAATTTTCGCATTTTTTGTAAGGGGTAACAT  
CAT

The match between the T2 sequence and the C1/C2 sequence is

30



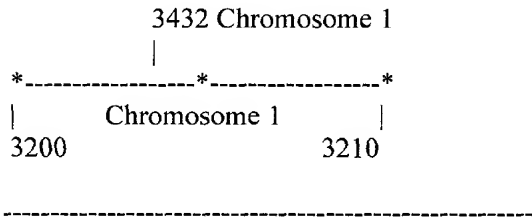


6. Permanent connectrons exist in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes.

C1/C2 short loops are normally expressed as the 3'UTR of some gene. A class of connectron relationships exist that permit one C1/C2 short loop to control the existence of one or more T1-T2 long loops without being subject to any expression controls other than those of the gene to which the C1/C2 is 3'UTR. These connectron relationships are described as "permanent". Permanent connectrons exist in prokaryotes, archea, single-celled eukaryotes and multi-celled eukaryotes.

Example of a prokaryote permanent connectron – E. coli

In this example the existence of the T1-T2 (3200-3210) long loop is controlled by a C1/C2 short loop (3432). The expression of this C1/C2 short loop is controlled only by the gene *btuB*.



A double stranded DNA loop of length 93.339 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3200. This T1 control element has the DNA sequence

AAGCGGCACTGCTCTTTAACAATTTATCAGACAATCTGTGTGGGCACTCG  
AAGATACGGATTCTTAACGTCGCAAGACGAAAAATGAATACCAAGTCTCA  
AGAGTGAACACGTAATTCATTACGAAGTTTAATTCTTTGAGCATCAAACCTT  
TTAAATTGAAGAGTTTGATCATGGCTCAGATTGAACGCTGGCGGCAGGCC  
TAACACATGCAAGTCGAACGGTAACAGGAAACAGCTTGCTGTTTCGCTGA  
CGAGTGGCGGACGGGTGAGTAATGTCTGGGAAACTGCCTGATGGAGGGG

GATAACTACTGGAAACGGTAGCTAATACCGCATAACGTCGCAAGACCAAA  
GAGGGGGACCTTCGGGCCTCTTGCCATC

This double stranded DNA loop is bounded on the right by a T2 control element  
whose identifier is 3310. This T2 control element has the DNA sequence

CAGACAATCTGTGTGGGCACTCGAAGATACGGATTCTTAACGTCGCAAGA  
CGAAAAATGAATACCAAGTCTCAAGAGTGAACACGTAATTCATTACGAAG  
TTTAATTCTTTGAGCGTCAAACCTTTTAAATTGAAGAGTTTGATCATGGCTC  
AGATTGAACGCTGGCGGCAGGCCTAACACATGCAAGTCGAACGGTAACA  
GGAAGAAGCTTGCTTCTTTGCTGACGAGTGGCGGACGGGTGAGTAATGTC  
TGGGAAACTGCCTGATGGAGGGGGATAACTACTGGAAACGGTAGCTAAT  
ACCGCATAACGTCGCAAGACCAAAGAGGGGGACCTTCGGGCCTCTTGCCA  
TCGGATGTGCCCAGATGGGATTAGCTAGT

This long T1/T2 double stranded DNA loop modulates the expression of the  
following genes

rrsC	gltU	rrlC	rrfC	aspT	trpT	yifA	yifE	yifB
ilvL	ilvG_1	ilvM	ilvE	ilvD	ilvA	ilvY	ilvC	ppiC
b3776	rep	gppA	rhlB	trxA	rhoL	rho	rfe	wzzE
wecB	rffH	wecD	wecE	wzxE	yifM_2	wecG	yifK	
argX	hisR	leuT	proM	aslB	aslA	hemY	hemX	
hemD	cyaA	cyaY	b3808	dapF	uvrD	b3814	corA	
yigF	yigG	rarD	yigl	pldA	recQ	yigJ	yigK	pldB
yigL	yigM	metR	metE	ysgA	udp	yigN	ubiE	yigP
b3836	yigU	yigW_1	rfaH	yigC	ubiB	fadA	fadB	
pepQ	trkH	hemG						

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2  
short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 3432 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene *btuB* and has the DNA sequence

5 TGC GCGGTCAGAAAATTATTTTAAATTTCTCTTGTCAGGCCGGAATAACT  
 CCCTATAATGCGCCACCACTGACACGGAACAACGGCAAACACGCCGCCGG  
 GTCAGCGGGGTTCTCCTGAGAACTCCGGCAGAGAAAGCAAAAATAAATG  
 CTTGACTCTGTAGCGGGAAGGCGTATTATGCACACC...TGCAACTCGACTC  
 CATGAAGTCGGAATCGCTAGTAATCGTGGATCAGAATGCCACGGTGAATA  
 10 CGTTCCCGGGCCTTGACACACCGCCCGTCACACCATGGGAGTGGGTTGC  
 AAAAGAAGTAGGTAGCTTAACCTTCGGGAGGGCGCTTACCACTTTGTGAT  
 TCATGACTGGGGTGAAGTCGTAACAAGGTAACCGTAGGGGAACCTGCGGT  
 TGGATCACCTCCTTACCTTAAAGAAGCGT

15 The match between the T1 sequence and the C1/C2 sequence is

AAGCGGCACTGCTCTTTAACAATTTATCAGACAATCTGTGTGGGCACTCG  
 AAGATACGGATTCTTAACGTCGCAAGACGAAAAATGAATACCAAGTCTCA  
 AGAGTGAACACGTAATTCATTACGAAGTTTAATTCTTTGAGC

20

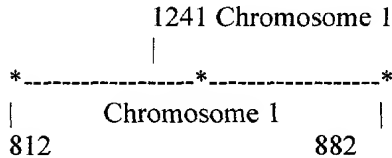
The match between the T2 sequence and the C1/C2 sequence is

CAGACAATCTGTGTGGGCACTCGAAGATACGGATTCTTAACGTCGCAAGA  
 CGAAAAATGAATACCAAGTCTCAAGAGTGAACACGTAATTCATTACGAAG  
 25 TTTAATTCTTTGAGCGTCAAACCTTTTAAATTGAAGAGTTTGATCATGGCTC  
 AGATTGAACGCTGGCGGCAGGCCTAACACATGCAAGTCGAACGGTAACA  
 GGAAGAAGCTTGCTTCTTTGCTGACGAGTGGCGGACGGGTGAGTAATGTC  
 TGGGAAACTGCCTGATGGAGGGGGATAACTACTGGAAACGGTAGCTAAT  
 ACCGCATAACGTCGCAAGACCAAAGAGGGGGACCTTCGGGCCTCTTGCCA  
 30 TCGGATGTGCCCAGATGGGATTAGCTAGT

-----

Example of an archea permanent connectron – H. pylori

In this example the existence of the T1-T2 (812-882) long loop is controlled by a C1/C2 short loop (1241). The expression of this C1/C2 short loop is controlled only by the gene HP1535.



A double stranded DNA loop of length 96.385 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 812. This T1 control element has the DNA sequence

TTTACTCATAGGGTTTTATAGTTCCTAGCGGAACTAAAGCA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 882. This T2 control element has the DNA sequence

TAGCGGAACTAAAGCATTCATCCCAAACACTAAAGATATTTGG

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

HP0999	HP1000	HP1001	HP1002	HP1003	HP1005	HP1006
HP1008	HP1009	HPtRNA-Pro	HP1010	HP1011	HP1013	HP1015
HP1017	HP1018	HP1020	HP1021	HP1022	HP1023	HP1024
HP1025	HP1027	HP1028	HP1030	HP1031	HP1033	HP1034
HP1038	HP1039	HP1040	HP1041	HP1042	HP1043	HP1044

HP1045    HP1046    HP1051    HP1052    HP1055    HP1056    HP1058  
 HP1060    HP1065    HPtRNA-Ser    HP1066    HP1067    HP1069    HP1070  
 HP1074    HP1075    HP1076    HP1077    HP1078    HP1079    HP1080  
 HP1081    HP1083    HP1084    HP1085    HP1088    HP1091    HP1092  
 5    HP1093    HP1094    HP1095    HP1096

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

10    A C1/C2 short loop on chromosome 1 whose identifier is 1241 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene HP1535 and has the DNA sequence

15    TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCATTTCATCCC  
 AAACA

The match between the T1 sequence and the C1/C2 sequence is

20    TTTTACTCATAGGGTTTTTATAGTTCCTAGCGGAACTAAAGCA

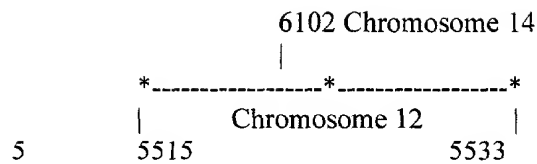
The match between the T2 sequence and the C1/C2 sequence is

TAGCGGAACTAAAGCATTTCATCCCAAACA

25    -----

Example of a single-celled permanent connectron – *S. cerevisiae*

30    In this example the existence of the T1-T2 (5515-5533) long loop is controlled by a C1/C2 short loop (6102). The expression of this C1/C2 short loop is controlled only by the gene YNL339C.



A double stranded DNA loop of length 6.466 kilo-bases on chromosome 12 is bounded on the left by a T1 sequence whose identifier is 5515. This T1 control element has the DNA sequence

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTTGTAGTGTGTTGCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTG

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 5533. This T2 control element has the DNA sequence

ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTTCTAGGGA  
ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTTAGTGTGTTGTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
TCCGGGTAAGAGACAACAGGGCT

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

YLR467W

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 14 whose identifier is 6102 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YNL339C and has the DNA sequence

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTTATTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTGAACATCCGGGTAAGAGACAACAGGGCT

The match between the T1 sequence and the C1/C2 sequence is

AGGAAATTGTTGTTACGAAAGTCAGTGATTATGTATTGTGTAGTATAGTAT  
ATTGTAAGAAATTTTTTTTCTAGGGAATATGCGTTTTGATGTAGTAGTAT  
TTCAGTGTGTTTATTAGTGTTTGTGTCACGGCAGTAGCGAGAGACAAGTG  
GGAAAGAGTAGGATAAAAAGACAATCTATAAAAAGTAAACATAAAATAA  
AGGTAGTAAGTAGCTTTTGGTTG

The match between the T2 sequence and the C1/C2 sequence is

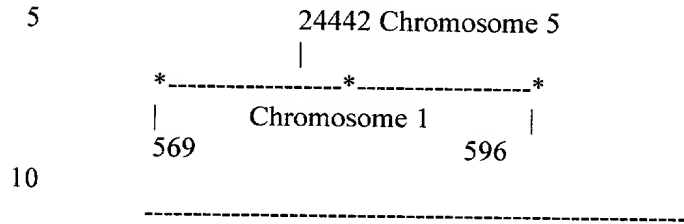
ATTATGTATTGTGTAGTATAGTATATTGTAAGAAATTTTTTTTCTAGGGA  
ATATGCGTTTTGATGTAGTAGTATTTCACTGTTTTGATTAGTGTTTGTGTC  
ACGGCAGTAGCGAGAGACAAGTGGGAAAGAGTAGGATAAAAAGACAATC  
TATAAAAAGTAAACATAAAATAAAGGTAGTAAGTAGCTTTTGGTTGAACA  
TCCGGGTAAGAGACAACAGGGCT

-----

Example of a multi-celled permanent connectron – *C. elegans*



In this example the existence of the T1-T2 (5515-5533) long loop is controlled by a C1/C2 short loop (6102). The expression of this C1/C2 short loop is controlled only by the gene YNL339C.



A double stranded DNA loop of length 30.606 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 569. This T1 control element has the DNA sequence

AAATCGAGCCCGTAAATCGACACAAGCGCTACAGTAGTC

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 596. This T2 control element has the DNA sequence

AGTGCTACAGTAGTCATTTAAGAATTACTGTAGTTTTTCGCT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 5 whose identifier is 24442 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene F20D6.4 and has the DNA sequence

GAGCCCGTAAATCGACACAAGCGCTACAGTAGTCATTTAAGAATTACTG  
TAGTTTTTC

The match between the T1 sequence and the C1/C2 sequence is

5

The match between the T2 sequence and the C1/C2 sequence is

GCTACAGTAGTCATTTAAAGAATTACTGTAGTTTTTC

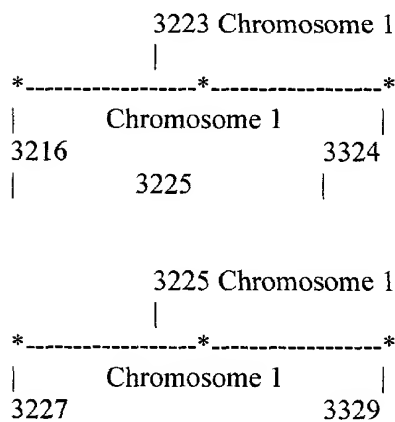
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7. Transient connectrons exist in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes.

A class of connectron relationships exist that permit one C1/C2 short loop to control the existence of one or more T1-T2 long loops such that this C1/C2 short loop is itself subject to expression control by another T1-T2 long loop which surrounds it. These connectron relationships are described as “transient”. Transient connectrons exist in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes.

Example of a prokaryote transient connectron – E. coli

In this example the existence of the T1-T2 (3227-3329) long loop is controlled by the C1/C2 (3225) short loop. The expression of this C1/C2 short loop is controlled by the existence of the T1-T2 (3216-3324) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the C1/C2 (3223) short loop. The C1/C2 (3225) short loop is the transient connectron.



A double stranded DNA loop of length 93.464 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3216. This T1 control element has the DNA sequence

AGCGCAAGCGAAGCTCTTGATCGAAGCCCCGGTAAACGGCGGCCGTA  
 ATAACGGTCCTAAGGTAGCGAAATTCCTTGTCGGGTAAGTTCCGACCTGC  
 ACGAATGGCGTAATGATGGCCAGGCTGTCTCCACCCGAGACTCAGTGAAA  
 TTGAACTCGCTGTGAAGATGCAGTGTACCCGCGGCAAGACGGAAAGACCC  
 5 CGTGAACCTTTACTATAGCTTGACACTGAACATTGAGCCTTGATGTGTAGG  
 ATAGGTGGGAGGCTTTGAAGTGTGGACGCCAGTCTGCATGGAGCCGACCT  
 TGAAATACCACCCTTTAATGTTTGATGTTCTAACGT

This double stranded DNA loop is bounded on the right by a T2 control element  
 whose identifier is 3324. This T2 control element has the DNA sequence

CCCGGTAAACGGCGGCCGTAACATAACGGTCCTAAGGTAGCGAAATTC  
 TTGTCGGGTAAGTTCCGACCTGCACGAATGGCGTAATGATGGCCAGGCTG  
 TCTCCACCCGAGACTCAGTGAAATTGAACTCGCTGTGAAGATGCAGTGTA  
 15 CCCGCGGCAAGACGGAAAGACCCCGTGAACCTTTACTATAGCTTGACACT  
 GAACATTGAGCCTTGATGTGTAGGATAGGTGGGAGGCTTTGAAGTGTGGA  
 CGCCAGTCTGCATGGAGCCGACCTTGAAATACCACCCTTTAATGTTTGATG  
 TTCTAACGTTGACCCGTAATCCGGGTTGCGGACAGT

This long T1/T2 double stranded DNA loop modulates the expression of the  
 following genes

rrfC	aspT	trpT	yifA	yifE	yifB	ilvL	ilvG_1	ilvM
ilvE	ilvD	ilvA	ilvY	ilvC	ppiC	b3776	rep	gppA
25 rhlB	trxA	rhoL	rho	rfe	wzzE	wecB	rffH	wecD
wecE	wzxE	yifM_2	wecG	yifK	argX	hisR	leuT	
proM	aslB	aslA	hemY	hemX	hemD	cyaA	cyaY	
b3808	dapF	uvrD	b3814	corA	yigF	yigG	rarD	yigI
pldA	recQ	yigJ	yigK	pldB	yigL	yigM	metR	metE
30 ysgA	udp	yigN	ubiE	yigP	b3836	yigU	yigW_1	rfaH
yigC	ubiB	fadA	fadB	pepQ	trkH	hemG	rrsA	ileT
rrlA								

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

5 A C1/C2 short loop on chromosome 1 whose identifier is 3225 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene rrlC and has the DNA sequence

10 AAACAGAATTTGCCTGGCGGCCGTAGCGCGGTGGTCCCACCTGACCCCAT  
GCCGAAGTCAGAAGTGAAACGCCGTAGCGCCGATGGTAGTGTGGGGTCTC  
CCCATGCGAGAGTAGGGAAGTCCAGGCATCAAATTA

15 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 3323 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene rrlA and has the DNA sequence

20 GCGAAGCTCTTGATCGAAGCCCCGGTAAACGGCGGCCGTAACTATAACGG  
TCCTAAGGTAGCGAAATTCCTTGTCGGGTAAAGTCCGACCTGCACGAATG  
GCGTAATGATGGCCAGGCTGTCTCCACCCGAGACTCAGTGAAATTGAACT  
CGCTGTGAAGATGCAGTGTACCCGCGGCAAGACGGA...AACAGAATTTGC  
25 CTGGCGGCAGTAGCGCGGTGGTCCCACCTGACCCCATGCCGAAGTCAGAA  
GTGAAACGCCGTAGCGCCGATGGTAGTGTGGGGTCTC

The match between the T1 sequence and the C1/C2 sequence is

30 GCGAAGCTCTTGATCGAAGCCCCGGTAAACGGCGGCCGTAACTATAACGG  
TCCTAAGGTAGCGAAATTCCTTGTCGGGTAAAGTCCGACCTGCACGAATG  
GCGTAATGATGGCCAGGCTGTCTCCACCCGAGACTCAGTGAAATTGAACT

CGCTGTGAAGATGCAGTGTACCCGCGGCAAGACGGAAAGACCCCGTGAA  
 CCTTTACTATAGCTTGACACTGAACATTGAGCCTTGATGTGTAGGATAGGT  
 GGGAGGCTTTGAAGTGTGGACGCCAGTCTGCATGGAGCCGACCTTGAAAT  
 ACCACCCTTTAATGTTTGATGTTCTAACGT

5

The match between the T2 sequence and the C1/C2 sequence is

CCCGGTAAACGGCGGCCGTAACCTATAACGGTCCTAAGGTAGCGAAATTCC  
 TTGTCGGGTAAGTTCCGACCTGCACGAATGGCGTAATGATGGCCAGGCTG  
 TCTCCACCCGAGACTCAGTGAAATTGAACTCGCTGTGAAGATGCAGTGTA  
 CCCGCGGCAAGACGGAAAGACCCCGTGAACCTTTACTATAGCTTGACACT  
 GAACATTGAGCCTTGATGTGTAGGATAGGTGGGAGGCTTTGAAGTGTGGA  
 CGCCAGTCTGCATGGAGCCGACCTTGAAATACCACCCTTTAATGTTTGATG  
 TTCTAACGTTGACCCGTAATCCGGGTTGCGGACAGT

10

15

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A double stranded DNA loop of length 93.749 kilo-bases on chromosome 1 is  
 bounded on the left by a T1 sequence whose identifier is 3227. This T1 control  
 element has the DNA sequence

AGCGCCGATGGTAGTGTGGGGTCTCCCCATGCGAGAGTAGGGAACTGCCA  
 GG

20

This double stranded DNA loop is bounded on the right by a T2 control element  
 whose identifier is 3329. This T2 control element has the DNA sequence

CATGCGAGAGTAGGGAACTGCCAGGCATCAAATAAAACGAAAGGCTCAG  
 TCG

25

30

This long T1/T2 double stranded DNA loop modulates the expression of the  
 following genes

	aspT	trpT	yifA	yifE	yifB	ilvL	ilvG_1	ilvM	ilvE
	ilvD	ilvA	ilvY	ilvC	ppiC	b3776	rep	gppA	rhlB
	trxA	rhoL	rho	rfe	wzzE	wecB	rffH	wecD	wecE
5	wzxE	yifM_2	wecG	yifK	argX	hisR	leuT	proM	
	aslB	aslA	hemY	hemX	hemD	cyaA	cyaY	b3808	
	dapF	uvrD	b3814	corA	yigF	yigG	rarD	yigI	pldA
	recQ	yigJ	yigK	pldb	yigL	yigM	metR	metE	ysgA
	udp	yigN	ubiE	yigP	b3836	yigU	yigW_1	rfaH	yigC
10	ubiB	fadA	fadB	pepQ	trkH	hemG	rrsA	ileT	rrlA
	rrfA								

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 3225 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene rrlC and has the DNA sequence

```
AAACAGAATTTGCCTGGCGGCCGTAGCGCGGTGGTCCCACCTGACCCCAT
GCCGAAGTCTCAGAAAGTCAAACGCGGTAGCGCCGATGGTAGTGTGGGGTCTC
CCCATGCGAGAGTAGGGAAGTCCAGGCATCAAATTA
```

The match between the T1 sequence and the C1/C2 sequence is

```
AGCGCCGATGGTAGTGTGGGGTCTCCCATGCGAGAGTAGGGAAGTCCCA
GG
```

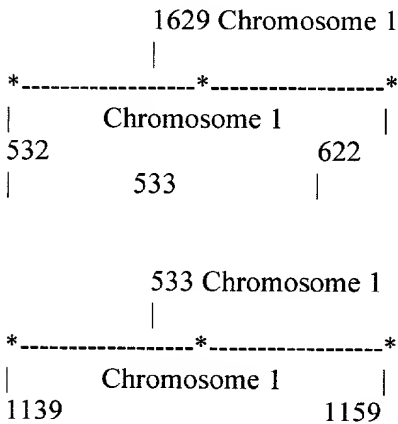
The match between the T2 sequence and the C1/C2 sequence is

```
CATGCGAGAGTAGGGAAGTCCAGGCATCAAAT
```

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Example of an archea transient connectron – *M. jannaschii*

In this example the existence of the T1-T2 (1139-1159) long loop is controlled by the C1/C2 (533) short loop. The expression of this C1/C2 short loop is controlled by the existence of the T1-T2 (532-622) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the C1/C2 (1629) short loop. The C1/C2 (533) short loop is the transient connectron.



A double stranded DNA loop of length 78.672 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 532. This T1 control element has the DNA sequence

ATATGTTTGAAATTTGAAAATAAGAGTATTAG

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 622. This T2 control element has the DNA sequence

TTGAAAATAAGAGCATTTAGAAGTTATTAATTAGTTCAAAGGATTTT



This long T1/T2 double stranded DNA loop modulates the expression of the following genes

	MJ0486	MJ0487	MJ0488	MJ0489	MJ0490	MJ0492	MJ0493
5	MJ0494	MJ0495	MJ0496	MJ0497	MJ0499	MJ0500	MJ0501
	MJ0502	MJ0503	MJ0504	MJ0506	MJ0507	MJ0508	MJ0509
	MJ0510	MJ0511	MJ0512	MJ0513	MJ0514	MJ0514	MJ0517
	MJ0519	MJ0520	MJ0521	MJ0522	MJ0523	MJ0525	MJ0526
	MJ0526	MJ0529	MJ0530	MJ0531	MJ0532	MJ0534	MJ0535
10	MJ0536	MJ0538	MJ0539	MJ0540	MJ0541	MJ0542	MJ0543
	MJ0544	MJ0545	MJ0547	MJ0548	MJ0549	MJ0550	MJ0552
	MJ0553	MJ0554	MJ0555	MJ0556	MJ0557	MJ0558	MJ0559
	MJ0560	MJ0561	MJ0562	MJ0563	MJ0564	MJ0565	MJ0566
	MJ0568	MJ0569	MJ0570				

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 1 whose identifier is 533 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ0485 and has the DNA sequence

ATTTTATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGA  
GTTTATTGAATT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 1629 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1597 and has the DNA sequence

ATATGTTTGAAATTTGAAAATAAGAGTATTTAGAAGTTATTAATTAGTTCA  
AAGGATTTTTATTTAATTTCTAAGGGTTTGCTGGTTTGATTATTTAGAATAT  
TTGAGTTTATTGAATTATTCAGATTTTTTAAAAATTA

5

The match between the T1 sequence and the C1/C2 sequence is

ATATGTTTGAAATTTGAAAATAAGAGTATTTAG

10

The match between the T2 sequence and the C1/C2 sequence is

ATTTAGAAGTTATTAATTAGTTCAAAGGATTTT

-----

15

A double stranded DNA loop of length 14.509 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 1139. This T1 control element has the DNA sequence

20

ATTTATTAATTAGTTCAAAGGATTTTTATTTAATTTCTAAGGGTTAGCTGG  
TTTGATTGTTTAAAATATTTGAGTTTA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1159. This T2 control element has the DNA sequence

25

ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
TGAATTATTCAGATTTTTTAAAAATTA

30

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

MJ1096 MJ1097 tRNA-Arg-3 MJ1098 MJ1099 MJ1100 MJ1101  
 MJ1102 MJ1103 MJ1104 MJ1105 MJ1106 MJ1107 MJ1108

5 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 533 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ0485 and has the DNA sequence

10 ATTTTATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGA  
 GTTTATTGAATT

The match between the T1 sequence and the C1/C2 sequence is

15 ATTTTATTTAATTTCTAAGGGTTAGCTGGTTTGATT

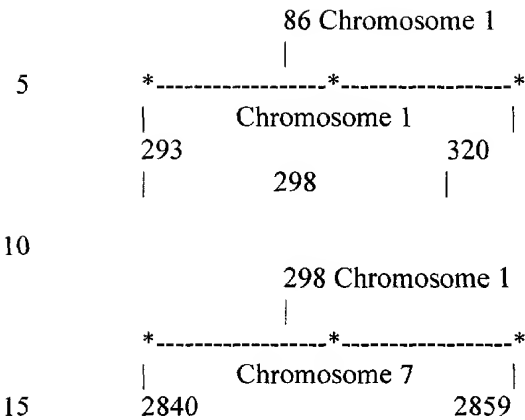
The match between the T2 sequence and the C1/C2 sequence is

20 ATTTAATTTCTAAGGGTTAGCTGGTTTGATTATTTAGAATATTTGAGTTTAT  
 TGAATT

-----

25 Example of a single-celled transient connectron – *S. cerevisiae*

In this example the existence of the T1-T2 (2840-2859) long loop is controlled by the C1/C2 (298) short loop. The expression of this C1/C2 short loop is controlled by the existence of the T1-T2 (293-320) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the C1/C2 (86) short loop. The C1/C2 (298) short loop is the transient connectron.



A double stranded DNA loop of length 38.470 kilo-bases on chromosome 2 is bounded on the left by a T1 sequence whose identifier is 293. This T1 control element has the DNA sequence

GAATTGTTGGAATAAAAATCCACTATCGTCTATCAACTAATAGTTATATTA  
TCAATATATTATCATATACGGTGTTAAGATGATGACATAAGTTATGAGAA  
GCTGTCATCGAAGTTAGAGGAAGCTGAAGTGCAAGGATTGATAATGTAAT  
AGGATAATGAAACATATAAAACGGAATGAGGAATAATCGTAATATTAGT  
ATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTTGAGGAGAAC  
TTCTAGT

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 320. This T2 control element has the DNA sequence

AATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTC  
GAGGAGAACTTCTAGTATATTCTGTA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

YBL005W-B TS(AGA)B YBL004W YBL003C YBL002W YBL001C  
 YBR001C YBR002C YBR003W YBR004C YBR005W YBR006W  
 YBR007C YBR008C YBR009C YBR010W YBR011C YBR012C

5

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 2 whose identifier is 298 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YBL005W-B and has the DNA sequence

10

ATCTATTACATTATGGGTGGTATGTTGGAATAAAAATCCACTATCGTCTAT  
 CAACTAATAGTTATATTATCAATATATTATCATATACGGTGTTAAGATGAT  
 GACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAAGTGCA  
 AGGATTGATAATGTAATAGGATAATGAAACATATAAAACGGAATGAGGA  
 ATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCT  
 ATATCCTTGAGGAGAACTTCTAGTATATTCTGTATACCTAATATTATAGCC  
 TTTATCAACAATGGAATCCCAACAATTATCTCAACATTC

15

20

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 86 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YAR009C and has the DNA sequence

25

ATCTATTACATTATGGGTGGTATGTTGGAATAGAAATCAACTATCATCTAC  
 TAACTAGTATTTACATTACTAGTATATTATCATATACGGTGTTAGAAGATG  
 ACGCAAATGATGAGAAATAGTCATCTAAATTAGTGGAAGCTGAAACGCA  
 AGGATTGATAATGTAATAGGATCAATGAATATAAAACATATAAAACGGAAT

30

GAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGG  
ATTCCTATATCCTCGAGGAGAACTTCTAGTATATTCTGTATACCTAATATT  
ATAGCCTTTATCAACAATGGAATCCCAACAATTATCTCAACATTCACCCAT  
TTCTCAGAA

5

The match between the T1 sequence and the C1/C2 sequence is

AAACATATAAAACGGAATGAGGAATAATCGTAATATTAGTATGTAGAAAT  
ATAGATTCCATTTTGAGGATTCCTATATCCT

10

The match between the T2 sequence and the C1/C2 sequence is

AATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTC  
GAGGAGAACTTCTAGTATATTCTGTA

15

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A double stranded DNA loop of length 5.302 kilo-bases on chromosome 7 is bounded  
on the left by a T1 sequence whose identifier is 2840. This T1 control element has  
the DNA sequence

20

TCTGTTGGAATAAAAATCCACTATCGTCTATCAACTAATAGTTATATTATC  
AATATATTATCATATACGGTGTTAAGATGATGACATAAGTTATGAGAAGC  
TGTCATCGAAGTTAGAGGAAGCTGAAACGCAAGGATTGATAATGTAATAG  
GATCAATGAATATAAACATATAAAACGGAATGAGGAATAATCGTAATATT  
AGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTCGAGGAG  
AACTTCTAGTATATTCTGTATACCTAAATTATAGCCTTTATCAACAATGGA  
ATCCCAACAA

25

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This double stranded DNA loop is bounded on the right by a T2 control element  
whose identifier is 2859. This T2 control element has the DNA sequence

CTATCAACTAATAGTTATATTATCAATATATTATCATATACGGTGTTAAGA  
 TGATGACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAAA  
 CGCAAGGATTGATAATGTAATAGGATCAATGAATATAAACATATAAAACG  
 GAATGAGGAATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTT  
 5 GAGGATTCCTATATCCTCGAGGAGAAGCTTCTAGTATATTCTGTATACCTAA  
 TATTATAGCCTTTATCAACAATGGAATCCCAACAATTATCTCAACATTCAC  
 ATATTTCTCAT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2  
 10 short loops.

A C1/C2 short loop on chromosome 2 whose identifier is 298 controls the expression  
 of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA  
 single strand that is 3'UTR to the gene YBL005W-B and has the DNA sequence

15 ATCTATTACATTATGGGTGGTATGTTGGAATAAAAATCCACTATCGTCTAT  
 CAACTAATAGTTATATTATCAATATATTATCATATACGGTGTTAAGATGAT  
 GACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAAGTGCA  
 AGGATTGATAATGTAATAGGATAATGAAACATATAAAACGGAATGAGGA  
 20 ATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCT  
 ATATCCTTGAGGAGAAGCTTCTAGTATATTCTGTATACCTAATATTATAGCC  
 TTTATCAACAATGGAATCCCAACAATTATCTCAACATTC

The match between the T1 sequence and the C1/C2 sequence is

25 TGTTGGAATAAAAATCCACTATCGTCTATCAACTAATAGTTATATTATCAA  
 TATTATCATATACGGTGTTAAGATGATGACATAAGTTATGAGAAGCTG  
 TCATCGAAGTTAGAGGAAGCTGAA

30 The match between the T2 sequence and the C1/C2 sequence is

CTATCAACTAATAGTTATATTATCAATATATTATCATATACGGTGTTAAGA  
TGATGACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAA

5

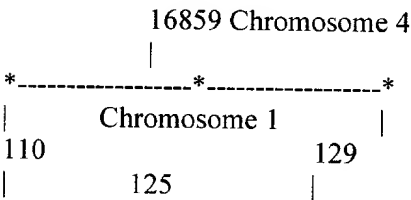
Example of a multi-celled transient connectron – *C. elegans*

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In this example the existence of the T1-T2 (22072-22108) long loop is controlled by the C1/C2 (125) short loop. The expression of this C1/C2 short loop is controlled by the existence of the T1-T2 (110-129) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the C1/C2 (16859) short loop. The C1/C2 (125) short loop is the transient connectron.

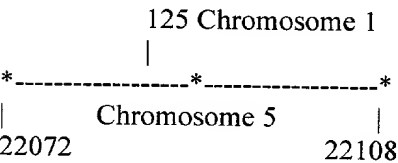
TGATGACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAA

15



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A double stranded DNA loop of length 18.855 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 110. This T1 control element has the DNA sequence

AGCTTAGGCTTAAGCTTAGGCTTAAGCTTAGGC

35

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 129. This T2 control element has the DNA sequence



TTCTCCCGCATTTTTTGTAGATCTACGTAGATCAAACCGAAATGAGGCACT  
TTCTGAATCCACGAGCTAGGCTTAAGCTTAGGCTTAAGCTTAGGCCTTTTC  
TCAGGCTTAGGCTTAGGCTTA

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This long T1/T2 double stranded DNA loop modulates the expression of the following genes

ZC123.3 ZC123.2

10

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 1 whose identifier is 125 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZC123.3 and has the DNA sequence

15

ACGCGCCGTAAATCTACCCCAGATATGGCCGAGCCAAAATGGCCTAGTTC  
GGCAAACCTCTTTCATTTCAATTTATGAGGGAAGCCAGAA

20

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

25

A C1/C2 short loop on chromosome 4 whose identifier is 16859 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene F58E2.7 and has the DNA sequence

CTTAGGCTTAAGCTTAGGCTTAAGCTTAGGCTTAAGCTTAGGCTTAAGCTT  
AGGCTTAAGCTTAGGCTTAAGCTTAGGCTTAAGCTTAGGCTTAAGCTTAG  
GCTTAAGCTTAGGCTTAAGCTTAGGCTTAAGCTTAGGCTTAAGCTTAGGCT  
TAAGCTTAGACTTA

30

The match between the T1 sequence and the C1/C2 sequence is

AGCTTAGGCTTAAGCTTAGGCTTAAGCTTAGGC

The match between the T2 sequence and the C1/C2 sequence is

TAGGCTTAAGCTTAGGCTTAAGCTTAGGC

A double stranded DNA loop of length 51.031 kilo-bases on chromosome 5 is bounded on the left by a T1 sequence whose identifier is 22072. This T1 control element has the DNA sequence

CGCAACGCGCCGTAAATCTACCCCAGATATGGCCGAGCCAAAATGACCTA  
GTTCGGC

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 22108. This T2 control element has the DNA sequence

TGACAATCGCCTGCCGGACAACGCGTGGAAAAGTGTCGTGTACTCCACAC  
GGACAAATACATTTAGTTTTACAATAAAATCGAACCGCGACGCGACACG  
CAACGCGACGTAAATCTACCCCAGATATGGCCGAGCCAAAATGGCCTAGT  
TCGGCAAACCTCTTCTATTTC

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

F36H9.3 F36H9.4 F36H9.5 F36H9.2 F36H9.1 F36H9.6

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 125 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZC123.3 and has the DNA sequence

ACGCGCCGTAAATCTACCCCAGATATGGCCGAGCCAAAATGGCCTAGTTC  
GGCAAACCTCTTTCAATTTATGAGGGAAGCCAGAA

The match between the T1 sequence and the C1/C2 sequence is

ACGCGCCGTAAATCTACCCCAGATATGGCCGAGCCAAAATG

The match between the T2 sequence and the C1/C2 sequence is

CGTAAATCTACCCCAGATATGGCCGAGCCAAAATGGCCTAGTTCGGCAAA  
CTCTT

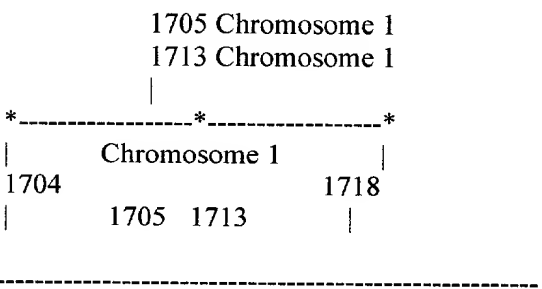
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8. Self-limiting connectrons occur in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes

A class of connectron relationships exist that permit one C1/C2 short loop to control the existence of the T1-T2 long loop that surrounds it. These connectron relationships are described as “self-limiting”. Self-limiting connectrons exist in prokaryotes, archaea, single-celled eukaryotes and multi-celled eukaryotes.

Example of a prokaryotic self-limiting connectrons – E. coli

In this example the existence of the T1-T2 (1704-1718) long loop is controlled by two C1/C2 (1705 and 1713) short loops. The expression of these C1/C2 short loops is controlled by the existence of the T1-T2 (1704-1718) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the two C1/C2 (1705 and 1713) short loops. The C1/C2 (1705 and 1713) short loops are the self-limiting connectrons.



A double stranded DNA loop of length 15.259 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 1704. This T1 control element has the DNA sequence

CGCCCCGTTACACGATTCCTCTGTAGTTCAGTCGGTAGAACGGCGGACT  
GTTAATCCGTATGTCACTGGT

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1718. This T2 control element has the DNA sequence

5 TTCAGTCGGTAGAACGGCGGACTGTTAATCCGTATGTCACTGGTTCGAGTC  
CAGTCAGAGGAGCCAAATTC

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

10 asnT b1978 b1979 b1980 shiA amn b1983 asnW  
yeeO asnU

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

15 A C1/C2 short loop on chromosome 1 whose identifier is 1705 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene and has the DNA sequence

20 CGCCCCGTTACACGATTCCTCTGTAGTTCAGTCGGTAGAACGGCGGACT  
GTTAATCCGTATGTCACTGGTTCGAGTCCAGTCAGAGGAGCCAAATTC

25 A C1/C2 short loop on chromosome 1 whose identifier is 1713 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene asnW and has the DNA sequence

30 CACGATTCCTCTGTAGTTCAGTCGGTAGAACGGCGGACTGTTAATCCGTAT  
GTCACTGGTTCGAGTCCAGTCAGAGGAGCCAAATT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 1705 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene and has the DNA sequence

CGCCCCGTTACACGATTCTCTGTAGTTCAGTCGGTAGAACGGCGGACT  
GTTAATCCGTATGTCACTGGTTCGAGTCCAGTCAGAGGAGCCAAATTC

The match between the T1 sequence and the C1/C2 sequence is

CGCCCCGTTACACGATTCTCTGTAGTTCAGTCGGTAGAACGGCGGACT  
GTTAATCCGTATGTCACTGGT

The match between the T2 sequence and the C1/C2 sequence is

TTCAGTCGGTAGAACGGCGGACTGTTAATCCGTATGTCACTGGTTCGAGTC  
CAGTCAGAGGAGCCAAATTC

A C1/C2 short loop on chromosome 1 whose identifier is 1713 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene asnW and has the DNA sequence

CACGATTCCTCTGTAGTTCAGTCGGTAGAACGGCGGACTGTTAATCCGTAT  
GTCACTGGTTCGAGTCCAGTCAGAGGAGCCAAATT

The match between the T1 sequence and the C1/C2 sequence is

CACGATTCCTCTGTAGTTCAGTCGGTAGAACGGCGGACTGTTAATCCGTAT  
GTCACTGGT

The match between the T2 sequence and the C1/C2 sequence is

TTCAGTCGGTAGAACGGCGGACTGTTAATCCGTATGTCACTGGTTCGAGTC  
CAGTCAGAGGAGCCAAATT

5

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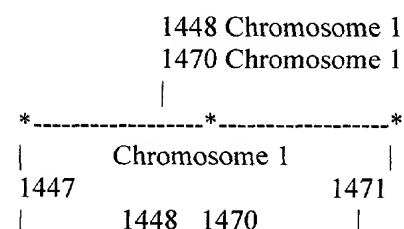
Example of a archea self-limiting connectrons – *M. jannaschii*

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In this example the existence of the T1-T2 (1447-1471) long loop is controlled by two C1/C2 (1448 and 1470) short loops. The expression of these C1/C2 short loops is controlled by the existence of the T1-T2 (1447-1471) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the two C1/C2 (1705 and 1713) short loops. The C1/C2 (1448 and 1470) short loops are the self-limiting connectrons.

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A double stranded DNA loop of length 22.675 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 1447. This T1 control element has the DNA sequence

30

TTATAGAACATTATGAAGCTTTTTACTCAACTAACAACCGTATCGAATTTA  
CCATTACTTGGAAATCTATTTAAACCTCTTTAATCTTATGATA

35

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1471. This T2 control element has the DNA sequence

CAACTAACAACCGTATCGAATTTACCATTACTTGGAATCTATTTAAAACC  
TCTTTAATCTTGTGATAATAAATTCTAATCGATTCGTGACTTAT

5 This long T1/T2 double stranded DNA loop modulates the expression of the following genes

MJ1402 MJ1403 MJ1404 MJ1405 MJ1406 MJ1407 MJ1408  
MJ1409 MJ1410 MJ1411 MJ1412 MJ1413 MJ1414 MJ1415  
10 MJ1416 MJ1417 MJ1418 MJ1419 MJ1420

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

15 A C1/C2 short loop on chromosome 1 whose identifier is 1448 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1401 and has the DNA sequence

20 TTATAGAACATTATGAAGCTTTTTACTCAACTAACAACCGTATCGAATTTA  
CCATTACTTGGAATCTATTTAAAACCTCTTTAATCTTATGATAATAAATT  
CTAATCGATTCGTGACTTAT

25 A C1/C2 short loop on chromosome 1 whose identifier is 1470 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1420 and has the DNA sequence

30 TTATAGAACATTATGAAGCTTTTTACTCAACTAACAACCGTATCGAATTTA  
CCATTACTTGGAATCTATTTAAAACCTCTTTAATCTTGTGATAATAAATT  
CTAATCGATTCGTG



The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 1470 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1420 and has the DNA sequence

TTATAGAACATTATGAAGCTTTTTACTCAACTAACAACCGTATCGAATTTA  
CCATTACTTGGAAATCTATTTAAAACCTCTTTAATCTTGTGATAATAAATT  
CTAATCGATTCTGTG

The match between the T1 sequence and the C1/C2 sequence is

TTATAGAACATTATGAAGCTTTTTACTCAACTAACAACCGTATCGAATTTA  
CCATTACTTGGAAATCTATTTAAAACCTCTTTAATCTT

The match between the T2 sequence and the C1/C2 sequence is

CAACTAACAACCGTATCGAATTTACCATTACTTGGAAATCTATTTAAAACC  
TCTTTAATCTTGTGATAATAAATTCTAATCGATTCTGTG

A C1/C2 short loop on chromosome 1 whose identifier is 1448 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene MJ1401 and has the DNA sequence

TTATAGAACATTATGAAGCTTTTTACTCAACTAACAACCGTATCGAATTTA  
CCATTACTTGGAAATCTATTTAAAACCTCTTTAATCTTATGATAATAAATT  
CTAATCGATTCTGTGACTTAT

The match between the T1 sequence and the C1/C2 sequence is

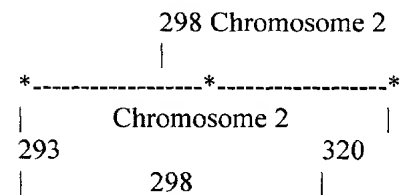
TTATAGAACATTATGAAGCTTTTTACTCAACTAACAACCGTATCGAATTTA  
CCATTACTTGGAATCTATTAAAACCTCTTTAATCTTATGATA

The match between the T2 sequence and the C1/C2 sequence is

CAACTAACAACCGTATCGAATTTACCATTACTTGGAATCTATTAAAACC  
TCTTTAATCTT

Example of a single-celled self-limiting connectron – *S. cerevisiae*

In this example the existence of the T1-T2 (293-320) long loop is controlled by C1/C2 (298) short loop. The expression of this C1/C2 short loop is controlled by the existence of the T1-T2 (293-320) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the C1/C2 (298) short loop. The C1/C2 (298) short loop is the self-limiting connectron.



A double stranded DNA loop of length 38.470 kilo-bases on chromosome 2 is bounded on the left by a T1 sequence whose identifier is 293. This T1 control element has the DNA sequence

GAATTGTTGGAATAAAAATCCACTATCGTCTATCAACTAATAGTTATATTA  
TCAATATATTATCATATACGGTGTTAAGATGATGACATAAGTTATGAGAA  
GCTGTCATCGAAGTTAGAGGAAGCTGAAGTGCAAGGATTGATAATGTAAT

AGGATAATGAAACATATAAAACGGAATGAGGAATAATCGTAATATTAGT  
 ATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTTGAGGAGAAC  
 TTCTAGT

5 This double stranded DNA loop is bounded on the right by a T2 control element  
 whose identifier is 320. This T2 control element has the DNA sequence

AATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCTC  
 GAGGAGAACTTCTAGTATATTCTGTA

10

This long T1/T2 double stranded DNA loop modulates the expression of the  
 following genes

YBL005W-B TS(AGA)B YBL004W YBL003C YBL002W YBL001C  
 YBR001C YBR002C YBR003W YBR004C YBR005W YBR006W  
 YBR007C YBR008C YBR009C YBR010W YBR011C YBR012C

This long T1/T2 double stranded DNA loop modulates the expression of the  
 following C1/C2 short loops

A C1/C2 short loop on chromosome 2 whose identifier is 298 controls the expression  
 of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is  
 expressed as a RNA single strand that is 3'UTR to the gene YBL005W-B and has the  
 DNA sequence

25

ATCTATTACATTATGGGTGGTATGTTGGAATAAAAATCCACTATCGTCTAT  
 CAACTAATAGTTATATTATCAATATATTATCATATACGGTGTTAAGATGAT  
 GACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAAGTGCA  
 AGGATTGATAATGTAATAGGATAATGAAACATATAAAACGGAATGAGGA  
 ATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCT  
 ATATCCTTGAGGAGAACTTCTAGTATATTCTGTATACCTAATATTATAGCC  
 TTTATCAACAATGGAATCCCAACAATTATCTCAACATTC

30

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

5 A C1/C2 short loop on chromosome 2 whose identifier is 298 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YBL005W-B and has the DNA sequence

10 ATCTATTACATTATGGGTGGTATGTTGGAATAAAAATCCACTATCGTCTAT  
CAACTAATAGTTATATTATCAATATATTATCATATACGGTGTTAAGATGAT  
GACATAAGTTATGAGAAGCTGTCATCGAAGTTAGAGGAAGCTGAAGTGCA  
AGGATTGATAATGTAATAGGATAATGAAACATATAAAACGGAATGAGGA  
ATAATCGTAATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCT  
15 ATATCCTTGAGGAGAACTTCTAGTATATTCTGTATACCTAATATTATAGCC  
TTTATCAACAATGGAATCCCAACAATTATCTCAACATTC

The match between the T1 sequence and the C1/C2 sequence is

20 TGTGGAATAAAAATCCACTATCGTCTATCAACTAATAGTTATATTATCAA  
TATATTATCATATACGGTGTTAAGATGATGACATAAGTTATGAGAAGCTG  
TCATCGAAGTTAGAGGAAGCTGAAGTGCAAGGATTGATAATGTAATAGGA  
TAATGAAACATATAAAACGGAATGAGGAATAATCGTAATATTAGTATGTA  
GAAATATAGATTCCATTTTGAGGATTCCTATATCCTTGAGGAGAACTTCTA  
GT

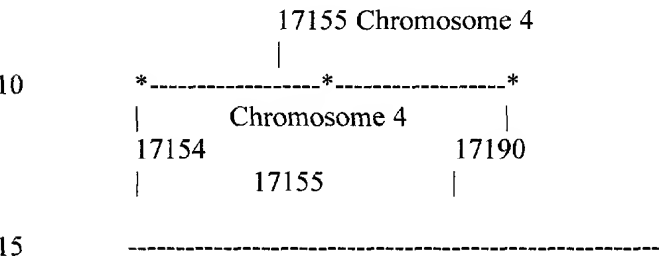
25 The match between the T2 sequence and the C1/C2 sequence is

AATATTAGTATGTAGAAATATAGATTCCATTTTGAGGATTCCTATATCCT

30 -----

Example of a multi-celled self-limiting connectron – *C. elegans*

In this example the existence of the T1-T2 (293-320) long loop is controlled by C1/C2 (298) short loop. The expression of this C1/C2 short loop is controlled by the existence of the T1-T2 (293-320) long loop. The existence of this T1-T2 long loop is itself determined by the expression of the C1/C2 (298) short loop. The C1/C2 (298) short loop is the self-limiting connectron.



A double stranded DNA loop of length 89,919 kilo-bases on chromosome 4 is bounded on the left by a T1 sequence whose identifier is 17154. This T1 control element has the DNA sequence

AAATTTCCGGCAAATCGGCAAACCTGGCAA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 17190. This T2 control element has the DNA sequence

AATTTGCCGATTTGCCGAATTTGTGCGACA

This long T1/T2 double stranded DNA loop modulates the expression of the following genes

R08C7.11 M01H9.2 M01H9.3 M01H9.4 M01H9.1 ZK180.1 ZK180.2  
ZK180.3 ZK180.4 ZK180.5 ZK180.6 ZK185.3 ZK185.2

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 4 whose identifier is 17155 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene R08C7.1 and has the DNA sequence

AAATTTCCGGCAAATCGGCAAACCTGGCAATTTGCCGATTTGCCGAATTTGT  
CGACA

A C1/C2 short loop on chromosome 4 whose identifier is 17171 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene ZK180.2 and has the DNA sequence

TGGAAATTTTCAAGAATTTCAATTTTAATCGGCAAAATTGTACGCATCCTATG  
AATTT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 4 whose identifier is 17155 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene R08C7.1 and has the DNA sequence

AAATTTCCGGCAAATCGGCAAACCTGGCAATTTGCCGATTTGCCGAATTTGT  
CGACA

The match between the T1 sequence and the C1/C2 sequence is

AAATTTCCGGCAAATCGGCAAACCTGGCAA

The match between the T2 sequence and the C1/C2 sequence is

AATTGCCGATTGCCGAATTGTCGACA

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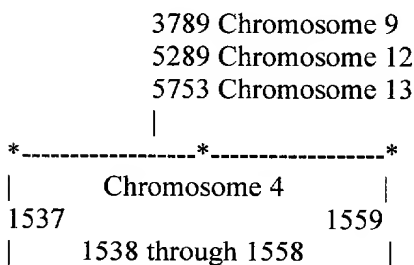
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9. Geneless connectrons exist in single-celled and multi-celled eukaryotes

Normally T1-T2 long loops contain genes whose expression is regulated by the existence of the long loop. When a T1-T2 long loop does not contain any genes it is described as being “geneless”. The existence of the T1-T2 long loop is itself controlled by one or more C1/C2 short loops that may be on the same or different chromosomes. The geneless T1-T2 long loops must contain one or more C1/C2 short loops.

Example of a single-celled geneless connectron – *S. cerevisiae*

In this example the existence of the T1-T2 (1537-1559) long loop is controlled by three C1/C2 (3789, 5289 and 5753) short loops. The expression of 21 C1/C2 (1538 through 1558) short loops are controlled by the existence of the T1-T2 (1537-1559) long loop.



A double stranded DNA loop of length 4.825 kilo-bases on chromosome 4 is bounded on the left by a T1 sequence whose identifier is 1537. This T1 control element has the DNA sequence

ATGAGATATATGTGGGTAATTAGATAATTGTTGGGATTCCATTGTTGATAA  
AGGCTATAATATTAGGTATACAGAATATACTAGAAGTTCTCCTCGAGGAT  
TTAGGAATCCATAAAAGGGAATCTGCAATTCTACACAATTCTATAAATAT  
TATTATCATCGTTTTATATGTTAATATTCATTGATCCTATTACATTATCAAT



CCTTGCGTTTCAGCTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGT  
CATCTTCTAACACCGTATATGATAATATACTAGTAACGTAAATACTAGTTA  
GTAGATGATAGTTGATTTTTATTCCAACATACCACCCATAATGTAATAGAT  
CTAAT

5

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1559. This T2 control element has the DNA sequence

ATGAGATATATGTGGGTAATTAGATAATTGTTGGGATTCCATTGTTGATAA  
AGGCTATAATATTAGGTATACAGAATATACTAGAAGTTCTCCTCGAGGAT  
TTAGGAATCCATAAAAAGGGAATCTGCAATTCTACACAATTCTATAAATAT  
TATTATCATCGTTTTATATGTTAATATTCATTGATCCTATTACATTATCAAT  
CCTTGCGTTTCAGCTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGT  
CATCTTCTAACACCGTATATGATAATATACTAGTAACGTAAATACTAGTTA  
GTAGATGATAGTTGATTTTTATTCCAACATACCACCCATAATGTAATAGAT  
CTAAT

10

15

There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 4 whose identifier is 1538 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

25

ATGAGATATATGTGGGTAATTAGATAATTGTTGGGATTCCATTGTTGATAA  
AGGCTATAATATTAGGTATACAGAATATACTAGAAGTTCTCCTCGAGGAT  
TTAGGAATCCATAAAAAGGGAATCTGCAATTCTACACAATTCTATAAATAT  
TATTATCATCGTTTTATATGTTAATATTCATTGATCCTATTACATTATCAAT  
CCTTGCGTTTCAGCTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGT  
CATCTTCTAACACCGTATATGATAATATACTAGTAACGTAAATACTAGTTA

30

GTAGATGATAGTTGATTTTTATTCCAACATACCACCCATAATGTAATAGAT  
CTAATGAATCCATTTGTTTGTTAATAGTTT

This T1-T2 loop also modulates the C1/C2 short loops numbered 1539 to 1557

A C1/C2 short loop on chromosome 4 whose identifier is 1558 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

AGCTTCTCATAACTTATGTCATCATCTTAACACCGTATATGATAATATATT  
GATAATATAACTTGTTGGAATAAAAATCAACTATCATCTACTAACTAGTAT  
TTACGTTACTAGTATATTATCATATACGGTGTTAGAAGATGACGCAAATG  
ATGAGAAATAGTCATCTAAATTAGTGGAAGCTGA...GTCTATCTGGCGAAT  
ATAAATTTTTACGCTACACACGTCATCGACATCTAAATATGACAGTCGCTG  
AACTGTTCTTAGATATCCATGCTATTTATGAAGAACAACAGGGATCGAGA  
AACAG

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 9 whose identifier is 3789 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YIL059C and has the DNA sequence

TTTATATGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCA  
GCTTCCACTAATTTAGATGACTATTTCTCATCTTTGCGTCATCTTCTAACA  
CCGTATATGATAATATACTAGTAACGTAAATACTAGTTAGTAGATGATAG  
TTGATTTTTATTCCAACAGTAT

The match between the T1 sequence and the C1/C2 sequence is

TTTATATGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCA  
 GCTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACA  
 CCGTATATGATAATATACTAGTAACGTAAATACTAGTTAGTAGATGATAG  
 TTGATTTTTATTCCAACA

5

The match between the T2 sequence and the C1/C2 sequence is

TTTATATGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCA  
 GCTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACA  
 CCGTATATGATAATATACTAGTAACGTAAATACTAGTTAGTAGATGATAG  
 TTGATTTTTATTCCAACA

10

A C1/C2 short loop on chromosome 12 whose identifier is 5289 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YLR301W and has the DNA sequence

15

GGTGAATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATA  
 TTAGGTATGTAGAATATACTAGAAGTTCTCCTCGAGGATTTAGGAATCCAT  
 AAAAGGGAATCTGCAATTCTACACAATTCTATAAATATTATTATCATCGTT  
 TTATATGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAG  
 CTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACAC  
 CGTATATGATAATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGT  
 TGATTTTTATTCCAACAC

25

The match between the T1 sequence and the C1/C2 sequence is

AGAATATACTAGAAGTTCTCCTCGAGGATTTAGGAATCCATAAAAGGGAA  
 TCTGCAATTCTACACAATTCTATAAATATTATTATCATCGTTTTATATGTTA  
 ATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAGCTTCCACTAA  
 TTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACACCGTATATGAT

30

AATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGTTGATTTTATT  
CCAACA

The match between the T2 sequence and the C1/C2 sequence is

5

AGGATTTAGGAATCCATAAAAAGGGAATCTGCAATTCTACACAATTCTATA  
AATATTATTATCATCGTTTTATATGTTAATATTCATTGATCCTATTACATTA  
TCAATCCTTGCGTTTCAGCTTCCACTAATTTAGATGACTATTTCTCATCATT  
TGCGTCATCTTCTAACACCGTATATGATAATATACTAGTAACGTAAATACT  
10 AGTTAGTAGATGATAGTTGATTTTATTCCAACA

A C1/C2 short loop on chromosome 13 whose identifier is 5753 controls the  
expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed  
as a RNA single strand that is 3'UTR to the gene YMR044W and has the DNA  
15 sequence

TTGAGAAATGGGGGAATGTTGAGATAATTGTTGGGATTCCATTGTTGATA  
AAGGCTATAATATTAGGTATACAGAATATACTAGAAGTTCTCCTCAAGGA  
TATAGGAATCCTCAAAATGGAATCTATATTTCTACATACTAATATTACGAT  
20 TATTCCTCATTCGTTTTATATGTTTCATTATCCTATTACATTATCAATCCT  
TGCACTTCAGCTTCCTCTAACTTCGATGACAGCTTCTCATAACTTATGTCA  
TCATCTTAACACCGTATATGATAATATATTGATAATATAACTATTAGTTGA  
TAGACGATAGTGGATTTTATTCCAACAT

25

The match between the T1 sequence and the C1/C2 sequence is

AGATAATTGTTGGGATTCCATTGTTGATAAAGGCTATAATATTAGGTATAC  
AGAATATACTAGAAGTTCTCCTC

30

The match between the T2 sequence and the C1/C2 sequence is

TTGTTGGGATTCCATTGTTGATAAAGGCTATAATATTAGGTATACAGAATA  
TACTAGAAGTTCTCCTCAAGGAT

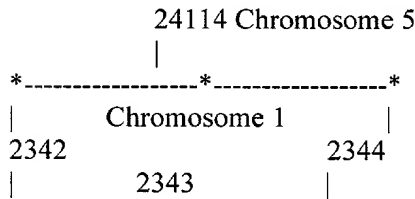
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5

Two examples of multi-celled geneless connectrons – *C. elegans*

In the first example the existence of the T1-T2 (2342-2344) long loop is controlled by the C1/C2 (24114) short loop. The expression of one C1/C2 (2343) short loop is controlled by the existence of the T1-T2 (2342-2344) long loop.

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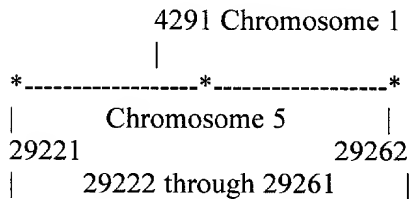


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In the second example the existence of the T1-T2 (29221-29262) long loop is controlled by the C1/C2 (24114) short loop. The expression of one C1/C2 (2343) short loop is controlled by the existence of the T1-T2 (2342-2344) long loop.

25



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A double stranded DNA loop of length 67.059 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 2342. This T1 control element has the DNA sequence

TGAAAACACTACAGTAATTCTTTAAATGACTACTGTAGC

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 2344. This T2 control element has the DNA sequence

CTACTGTAGCGCTTGTGTCGATTACGGGCTCGATT

There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 1 whose identifier is 2343 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

TCGACACAAGCGCTACAGTAGCTATTTAAAGAATTACTGTAGTTTTTCGCTACGAGATATTT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 5 whose identifier is 24114 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene C13F10.5 and has the DNA sequence

GCGAAAACACTACAGTAATTCTTTAAATGACTACTGTAGCGCTTGTGTCGATTACGGGCTCGATTTTCG

The match between the T1 sequence and the C1/C2 sequence is

GAAAACTACAGTAATTCTTTAAATGACTACTGTAGC

The match between the T2 sequence and the C1/C2 sequence is

CTACTGTAGCGCTTGTGTCGATTACGGGCTCGATT

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A double stranded DNA loop of length 41.297 kilo-bases on chromosome 5 is bounded on the left by a T1 sequence whose identifier is 29221. This T1 control element has the DNA sequence

TTTAAATTTCCCGCCAAAAATTGACTGAAAATTTGGATTTTCTTTCCAAAA  
ATTGACAGAAA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 29262. This T2 control element has the DNA sequence

TGAAAATTTGAATTTCCCGCCAAAAATTAAC

There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 5 whose identifier is 29222 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

AATTTCCCGCCAAAAATTGACTGAAAATTTGGATTTTCTTTCCAAAAATTG  
ACAGAAA

This T1-T2 loop also modulates the C1/C2 short loops numbered 29223 to 29260

A C1/C2 short loop on chromosome 5 whose identifier is 29261 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

AAAATTGACTGAAAATTTGAATTTCCAGCCAAAAATTGACTGAAAATTTG  
AATT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 1 whose identifier is 4291 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene Y43F8C.5 and has the DNA sequence

AAAATTAAGTAAAATTTGAATTTCCCGCCAAAAATTGACTGAAAATTTG  
AATTTCCCGCCAAAAAAATTGACTGAAAATTTGAATTTCCCGCCAAAA  
TTGACTGAAAATTTGAATTTCCCGCCAAAAATTAATTGAAAATTTGAATTT  
CCCGCCAAAAATTAATTGAACTTTGAATTTTCAA...ATTTCCCGCCAAAA  
ATTAATTGAACTTTGAATTTTCAAATTTCCCGCCAAAAATTGACTGAAAA  
TTTGAATTTCCCGCCAAAAATTAATTGAAAATTTGAATTTTGAATTTCC  
GCCAAAAATGACTGA

The match between the T1 sequence and the C1/C2 sequence is

TTTAAATTTCCCGCCAAAAATTGACTGAAAATTTG

The match between the T2 sequence and the C1/C2 sequence is



AAAAAAATTGACTGAAAATTTGAATTTCCCGCCAAAAATTGA

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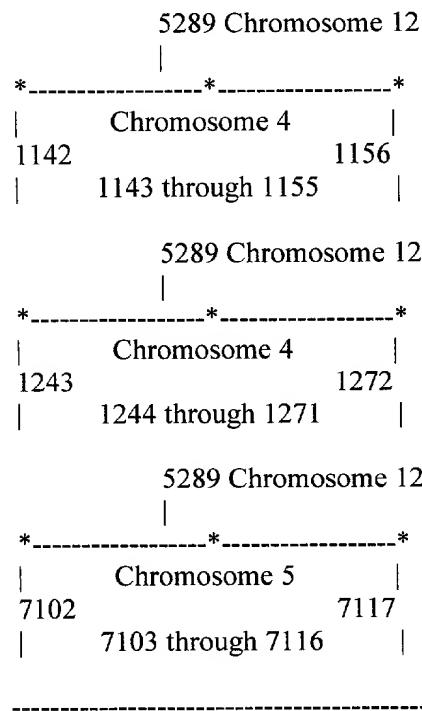
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10. One connectron controls many geneless connectrons in single-celled and multi-celled eukaryotes

One C1/C2 short loop can control the existence of many geneless T1-T2 long loops.

Example of a single-celled geneless connectron – *S. cerevisiae*

In this example the existence of the three T1-T2 (1142-1156, 1242-1272 and 7102-7117) long loops is controlled by the C1/C2 (5289) short loop.



A double stranded DNA loop of length 5.337 kilo-bases on chromosome 4 is bounded on the left by a T1 sequence whose identifier is 1142. This T1 control element has the DNA sequence

ATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATATTAGG  
TATGTAGATATACTAGAAGTTCTCCTCGAGGATTTAGGAATCCATAAAAG

GGAATCTGCAATTCTACACAATTCTATAAATATTATTATCATCATTTTATA  
 TGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAGCTTCC  
 ACTAATTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACACCGTAT  
 ATGATAATATACTAGTAACGTAATACTAGTTAGTAGATGATAGTTGATTT  
 TTATTCCAACA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1156. This T2 control element has the DNA sequence

TTTTAATAAGGCAATAATATTAGGTATGTAGATATACTAGAAGTTCTCCTC  
 CAGGATTTAGGAATCCATAAAAGGGAATCTGCAATTCTACACAATTCTAT  
 AAATATTATTATCATCATTTTATATGTTAATATTCATTGATCCTATTACATT  
 ATCAATCCTTGCGTTTCAGCTTCCACTAATTTAGATGACTATTTCTCATCAT  
 TTGCGTCATCTTCTAACACCGTATATGATAATATACTAGTAACGTAATACT  
 TAGTTAGTAGATGATAGTTGATTTTTATTCCAACAAGAA

There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 4 whose identifier is 1143 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

ATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATATTAGG  
 TATGTAGATATACTAGAAGTTCTCCTCGAGGATTTAGGAATCCATAAAAG  
 GGAATCTGCAATTCTACACAATTCTATAAATATTATTATCATCATTTTATA  
 TGTTAATATTCATTGATCCTATTACATTATCAAT...CTCTAAGTCTCATTGCC  
 TTTGTGCCAAAAAATCTGTTTCTAAATTTCTCTTCATTTGTAGACTTAATTA  
 TACTGATCGTTGATCTACTATCAGTAAGTAAGCCTTTAATAATTGGTTTCT  
 TGTTAAGTTCTTGACAAGGTGACTGAGGTTATTCAATAGCGG

This T1-T2 loop also modulates the C1/C2 short loops numbered 1144 to 1154

A C1/C2 short loop on chromosome 4 whose identifier is 1155 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

GAGGAGAACTTCTAGTATATCTACATACCTAATATTATTGCCTTATTAAAA  
ATGGAATCCCAACAATTA

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 12 whose identifier is 5289 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YLR301W and has the DNA sequence

GGTGAATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATA  
TTAGGTATGTAGAATATACTAGAAAGTTCTCCTCGAGGATTAGGAATCCAT  
AAAAGGGAATCTGCAATTCTACACAATTCTATAAATATTATTATCATCGTT  
TTATATGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAG  
CTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACAC  
CGTATATGATAATATACTAGTACGTAAATACTAGTTAGTAGATGATAGTT  
GATTTTTATTCCAACAC

The match between the T1 sequence and the C1/C2 sequence is

ATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATATTAGG  
TATGTAGA

The match between the T2 sequence and the C1/C2 sequence is

TTTTAATAAGGCAATAATATTAGGTATGTAGA

A double stranded DNA loop of length 5.251 kilo-bases on chromosome 4 is bounded on the left by a T1 sequence whose identifier is 1243. This T1 control element has the DNA sequence

CGTGTTTTATCTCATGTTGTTTCGTTTTGTTATTGAGATATATGTGGGTAATT  
AGATAATTGTTGGGATTCCATTGTTGATAAAGGCTATAATATTAGGTATAC  
AGAATATACTAGAAGTTCTCCTCGAGGATTTAGGAATCCATAAAAGGGAA  
TCTGCAATTCTACACAATTCTATAAAATATTATTATCATCGTTTTATATGTTA  
ATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAGCTTCCACTAA  
TTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACACCGTATATGAT  
AATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGTTGATTTTTATT  
CCAACA

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 1272. This T2 control element has the DNA sequence

TGAGATATATGTGGGTAATTAGATAATTGTTGGGATTCCATTGTTGATAAA  
GGCTATAATATTAGGTATACAGAATATACTAGAAGTTCTCCTCGAGGATTT  
AGGAATCCATAAAAGGGAATCTGCAATTCTACACAATTCTATAAAATATTA  
TTATCATCGTTTTATATGTTAATATTCATTGATC...TATACTAGTAACGTAA  
ATACTAGTTAGTAGATGATAGTTGATTTTTATTCCAACAGTTATAAGGTTG  
TTTCATATGTGTTTTATGAA

There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 4 whose identifier is 1244 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

5

TTTATCTCATGTTGTTTCGTTTTGTTATTGAGATATATGTGGGTAATTAGATA  
ATTGTTGGGATTCCATTGTTGATAAAGGCTATAATATTAGGTATACAGAAT  
ATACTAGAAGTTCTCCTCGAGGATTTAGGAATCCATAAAAGGGAATCTGC  
AATTCTACACAATTCTATAAATATTATTATCAT...GTCTCGATGTAGTATAC  
10 GTATAAATTATTACCTGATACTTCATCTCTAAGTCTCATTGCCTTTGTGCCA  
AAAAATCTGTTTCTAAATTTCTCTTCATTTGTAGACTTAATTATACTGATCG  
TTGATCTACTATCAGTAAGT

This T1-T2 loop also modulates the C1/C2 short loops numbered 1245 to 1270

15

A C1/C2 short loop on chromosome 4 whose identifier is 1271 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

20

TGTTGTATCTCAAAATGAGATATGTCAGTATGACAATACGTCATCCTAAAC  
GTTCATAAAACACATATGAAACAACCTTATAACTGTTGGAATAAAAATCA  
ACTATCATCTACTAACTAGTATTTACGTTACTAGTATATTATCATATACGG  
TGTTAGAAGATGACGCAAATGATGAGAAATAGTC...CAACAATGGAATCC  
CAACAATTATCTAATTACCCACATATATCTCATGGTAGCGCCTGTGCTTCG  
25 GTTACTTCTAAGGAAGTCCACACAAATCAAGATCCGTTAGACGTTTCAGC  
TTCCAAAA

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

30

A C1/C2 short loop on chromosome 12 whose identifier is 5289 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed

as a RNA single strand that is 3'UTR to the gene YLR301W and has the DNA sequence

5 GGTGAATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATA  
TTAGGTATGTAGAATATACTAGAAAGTTCTCCTCGAGGATTTAGGAATCCAT  
AAAAGGGAATCTGCAATTCTACACAATTCTATAAAATATTATTATCATCGTT  
TTATATGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAG  
CTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACAC  
CGTATATGATAATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGT  
10 TGATTTTTATTCCAACAC

The match between the T1 sequence and the C1/C2 sequence is

15 AGAATATACTAGAAAGTTCTCCTCGAGGATTAGGAATCCATAAAAGGGAA  
TCTGCAATTCTACACAATTCTATAAAATATTATTATCATCGTTTTATATGTTA  
ATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAGCTTCCACTAA  
TTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACACCGTATATGAT  
AATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGTTGATTTTTATT  
CCAACA

The match between the T2 sequence and the C1/C2 sequence is

25 AGAATATACTAGAAAGTTCTCCTCGAGGATTAGGAATCCATAAAAGGGAA  
TCTGCAATTCTACACAATTCTATAAAATATTATTATCATCGTTTTATATGTTA  
ATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAGCTTCCACTAA  
TTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACACCGTATATGAT  
AATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGTTGATTTTTATT  
CCAACA

30 -----

A double stranded DNA loop of length 5.296 kilo-bases on chromosome 15 is bounded on the left

by a T1 sequence whose identifier is 7102. This T1 control element has the DNA sequence

5

CATGATTAATATGACCAATCGGCGTGTGTTTTGAAAAGTGGGTGAATTTT  
GAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATATTAGGTATGT  
AGAATGTACTAGAAGTTCTCCTCAAGGATTTAGGAATCCATGAAAGGGAA  
TCTGCAATTCTACACAATTCTATAAATATTATTATCATCATTTTATATGTTA  
10 ATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAGCTTCCACTAA  
TTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACACCGTATATGAT  
AATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGTTGATTTTTATT  
CCAACA

15

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 7117. This T2 control element has the DNA sequence

20

TGAAAAGTGGGTGAATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAG  
GCAATAATATTAGGTATGTAGAATGTACTAGAAGTTCTCCTCAAGGATTT  
20 AGGAATCCATGAAAGGGAATCTGCAATTCTACACAATTCTATAAATATTA  
TTATCATCATTTTATATGTTAATATTCATTGATCCTATTACATTATCAATCC  
TTGCGTTTCAGCTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGTCA  
TCTTCTAACACCGTATATGATAATACTAGTAACGTAAATACTAGTTAGT  
AGATGATAGTTGATTTTTATTCCAACAGTTTTATATACCTCTCTTATTTAGT  
25 ATAAGAA

There are no genes controlled by this T1/T2 loop.

30

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops



A C1/C2 short loop on chromosome 15 whose identifier is 7103 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

5 AAGAACATTGCTGATGTGATGACAAAACCTCTTCCGATAAAAACATTTAA  
ACTATTAATAACAAATGGATTCATTAGATCTATTACATTATGGGTGGTAT  
GTTGGAATAAAAATCAACTATCATCTACTAACTAGTATTTACGTTACTAGT  
ATATTATCATATACGGTGTTAGAAGATGACGCAAATGATGAGAAATAGTC  
ATCTAAATTAGTGGAAGCTGAAACGCAAGGATTGATAATGTAATAGGATC  
10 AATGAATATTAACATATAAAATGATGATAATAATATTTATAGAATTGTGT  
AGAATTGCAGATTCCCTTTTCATGGATTCTTAAATCCTTGAGGAGAACTTCT  
AGTA

This T1-T2 loop also modulates the C1/C2 short loops numbered 7104 to 7115

15 A C1/C2 short loop on chromosome 15 whose identifier is 7116 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

20 CCATTCTGTGGAGGTGGTACTGAAGCAGGTTGAGGAGAGACATGATGATG  
GTTCTCTGGAACAGCT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

25 A C1/C2 short loop on chromosome 12 whose identifier is 5289 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene YLR301W and has the DNA sequence

30 GGTGAATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATA  
TTAGGTATGTAGAATATACTAGAAGTTCTCCTCGAGGATTAGGAATCCAT

AAAAGGGAATCTGCAATTCTACACAATTCTATAAATATTATTATCATCGTT  
 TTATATGTTAATATTCATTGATCCTATTACATTATCAATCCTTGCGTTTCAG  
 CTTCCACTAATTTAGATGACTATTTCTCATCATTTGCGTCATCTTCTAACAC  
 CGTATATGATAATATACTAGTAACGTAAATACTAGTTAGTAGATGATAGT  
 5 TGATTTTTATTCCAACAC

The match between the T1 sequence and the C1/C2 sequence is

GGTGAATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATA  
 10 TTAGGTATGTAGAAT

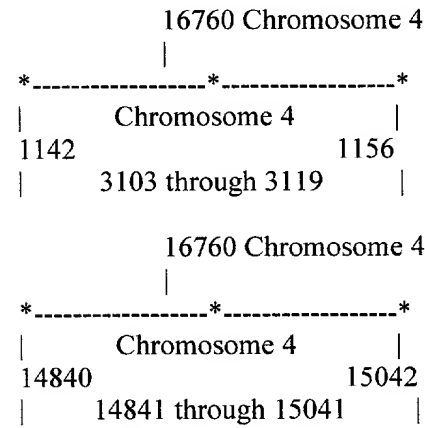
The match between the T2 sequence and the C1/C2 sequence is

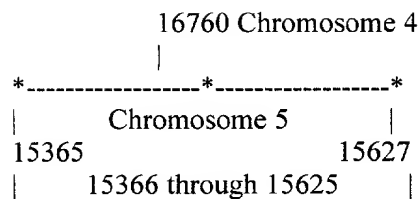
GGTGAATTTTGAGATAATTGTTGGGATTCCATTTTAAATAAGGCAATAATA  
 15 TTAGGTATGTAGAAT

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Example of a multi-celled geneless connectron – *C. elegans*

In this example the existence of the three T1-T2 (1142-1156, 14840-15042 and  
 15365-15627) long loops is controlled by the C1/C2 (16760) short loop.





A double stranded DNA loop of length 15.894 kilo-bases on chromosome 1 is bounded on the left by a T1 sequence whose identifier is 3101. This T1 control element has the DNA sequence

CAAATCGGCAAATTGCCGGAATTGAACATTTC

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 3120. This T2 control element has the DNA sequence

AAACGATTTTCCGGCAAATCGGCAAATTGCCGGAATTGTAATTTCCGGC  
AAAT

There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

A C1/C2 short loop on chromosome 1 whose identifier is 3103 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

TTAAAATTTCCGGCAAATCGGCAAATTGGCAGAAATGAAACTCACGGCAA  
ATCGG

This T1-T2 loop also modulates the C1/C2 short loops numbered 3104 to 3118

A C1/C2 short loop on chromosome 1 whose identifier is 3119 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

CCCGCATTTTTTTGTAGATCAAACCGTAATGGGACGGCCTGGCAACACGTG  
ATTTTCCAAAT

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 4 whose identifier is 16760 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene T23E1.2 and has the DNA sequence

GGCAAATTGCCGAAATTGAACATTTCCGGCAAATCGGCAAATTGCCGGAA  
TTGAACATTTCCGGCAAATCGGCAAATTGCCGGAATTGAACATTTCCGGC  
AAATCGGCAAATTGCCGGAATTGA

The match between the T1 sequence and the C1/C2 sequence is

CAAATCGGCAAATTGCCGGAATTGAACATTTCC

The match between the T2 sequence and the C1/C2 sequence is

TTTCCGGCAAATCGGCAAATTGCCGGAATTG

-----

A double stranded DNA loop of length 86.977 kilo-bases on chromosome 3 is bounded on the left by a T1 sequence whose identifier is 14840. This T1 control element has the DNA sequence

AAAAATTTCCGGCAAGTCGGCAATTTTCCGAAAATGAAAATTTCCGGCAA  
ATCGGCAAATTGCCGGAATTGAAAATTCCTGGCAAATCAGCAAATTTGCG  
GCAAATCGGCAATTTGCCGAAAATGAAAATTTCCGGCAAAT

5

This double stranded DNA loop is bounded on the right by a T2 control element whose identifier is 15042. This T2 control element has the DNA sequence

CAAATCGGTAGGTAAATTGGCCAAACTTGAAAATTTCCGGCAAATCGGCA  
AATTCCGCGAACTGAACATTTCCGGCAAATCGGCAAATTGCTCGAACT

10

There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the following C1/C2 short loops

15

A C1/C2 short loop on chromosome 3 whose identifier is 14841 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

AAAAATTTCCGGCAAGTCGGCAATTTTCCGAAAATGAAAATTTCCGGCAA  
ATCGGCAAATTGCCGGAATTGAAAATTCCTGGCAAATCAGCAAATTTGCG  
GCAAATCGGCAATTTGCCGAAAATGAAAATTTCCGGCAAAT

20

25

This T1-T2 loop also modulates the C1/C2 short loops numbered 14842 to 15040

A C1/C2 short loop on chromosome 3 whose identifier is 15041 controls the expression of the genes of one or more other T1/T2 long loops. This C1/C2 short loop has the DNA sequence

CGGCAATTGCCGTTCCGGCAATTTGCCAATTTGCCGGAATTTTCAATTCCG  
GCAA

30

The expression of genes in this T1/T2 long loop is controlled by the following C1/C2 short loops.

A C1/C2 short loop on chromosome 4 whose identifier is 16760 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene T23E1.2 and has the DNA sequence

```
GGCAAATTGCCGAAATTGAACATTTCCGGCAAATCGGCAAATTGCCGGAA
TTGAACATTTCCGGCAAATCGGCAAATTGCCGGAATTGAACATTTCCGGC
AAATCGGCAAATTGCCGGAATTGA
```

The match between the T1 sequence and the C1/C2 sequence is

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ATTTCCGGCAAATCGGCAAATTGCCGGAATTGAA
```

The match between the T2 sequence and the C1/C2 sequence is

```
TGAACATTTCCGGCAAATCGGCAAATTGC
```

A double stranded DNA loop of length 98.488 kilo-bases on chromosome 3 is bounded on the left by a T1 sequence whose identifier is 15365. This T1 control element has the DNA sequence

```
AAAATTTCCGGCAAATCGGCAATTTGCCAAAAATTGAAATTTCCGGCAAA
TCGGCAATTTGTCAAAAATGAAAATTTCCGGCAAATCGGCAAATTGCCGA
AAATGAAAATTTCCGGCAAATCGGCAAATTTCCGGAATGAAAATTTCCG
GCAAATCGGCAATTTGCCATAAATGAACATTTCCGG...GGCGAAAATTAAA
ATTTCCGCCATATCGGCAATTTGCCAAAAAATTAAAATTTCCGGCAAATC
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GGCAAATTGCCGGAATTCAAAATTTCCGGCAAACCGGCAAATTGCCGGAA  
CTCAAAATTTCCCGGCAAATCAGCAAATTGCCGGAATT

5 This double stranded DNA loop is bounded on the right by a T2 control element  
whose identifier is 15627. This T2 control element has the DNA sequence

TGGCAAACCGGCAAATTGCCGGAATTGAACATTTCCGGCAAATCGGCAAT  
TTGCCGGAATTGAAATTT

10 There are no genes controlled by this T1/T2 loop.

This long T1/T2 double stranded DNA loop modulates the expression of the  
following C1/C2 short loops

15 A C1/C2 short loop on chromosome 3 whose identifier is 15366 controls the  
expression of the genes of one or more other T1/T2 long loops. This C1/C2 short  
loop has the DNA sequence

20 TGCCGATTTGCCGGAAATTTTCATTTTCGGCAATTTGCCGATTTGCCGGAA  
ATTTTCATT

This T1-T2 loop also modulates the C1/C2 short loops numbered 15366 to 15624

25 A C1/C2 short loop on chromosome 3 whose identifier is 15625 controls the  
expression of the genes of one or more other T1/T2 long loops. This C1/C2 short  
loop has the DNA sequence

TCAAGCAAATTGTCAAATTCGCGGAATAAACATTTCCGGCAAATCGGCA  
AATT

30 The expression of genes in this T1/T2 long loop is controlled by the following C1/C2  
short loops.

A C1/C2 short loop on chromosome 4 whose identifier is 16760 controls the expression of the genes in this T1/T2 long loop. This C1/C2 short loop is expressed as a RNA single strand that is 3'UTR to the gene T23E1.2 and has the DNA sequence

5

GGCAAATTGCCGAAATTGAACATTTCCGGCAAATCGGCAAATTGCCGGAA  
TTGAACATTTCCGGCAAATCGGCAAATTGCCGGAATTGAACATTTCCGGC  
AAATCGGCAAATTGCCGGAATTGA

10

The match between the T1 sequence and the C1/C2 sequence is

ATTTCCGGCAAATCGGCAAATTGCCGGAATT

The match between the T2 sequence and the C1/C2 sequence is

15

CGGCAAATTGCCGGAATTGAACATTTCCGGCAAATCGGCAA

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